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#### INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification <sup>6</sup> :	1	11) International Publication Number: WO 99/21890
C07K 14/705	A1	43) International Publication Date: 6 May 1999 (06.05.99
(21) International Application Number: PCT/SI (22) International Filing Date: 27 October 1998  (30) Priority Data: 9703914-3 27 October 1997 (27.10.97 9800864-2 16 March 1998 (16.03.98) 9802575-2 17 July 1998 (17.07.98)  (71) Applicant (for all designated States except US): AKTIEBOLAG [SE/SE]; S-151 85 Södertälje (SC)  (72) Inventor; and (75) Inventor/Applicant (for US only): EKSTRAN [SE/SE]; Astra Hässle AB, Tvistevägen 48, Umeå (SE).  (74) Agent: ASTRA AKTIEBOLAG; Patent Dept., Södertälje (SE).	ASTE E). ID, Jon S-907	BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KF, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW Eurasian patent (AT, BE, CH, CY, DE, DK, ES, FI, FF GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BI BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN TD, TG).  Published  With international search report.  Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt amendments.

(57) Abstract

The present invention relates to a nucleic acid molecule encoding a GABA<sub>B</sub> receptor, or a functionally equivalent modified form thereof, said receptor being selected from the group consisting of human and canine GABA<sub>B</sub> receptors.

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#### TECHNICAL FIELD

The present invention relates in particular to nucleic acid molecules encoding GABA<sub>B</sub> receptors, and to methods for screening for compounds which are inhibitors of transient lower esophageal sphincter relaxations (TLESR), said methods comprising the use of a nucleic acid molecule encoding a GABA<sub>B</sub> receptor.

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#### **BACKGROUND ART**

#### GABA<sub>B</sub> receptors

- 15 GABA (4-aminobutanoic acid) is an endogenous neurotransmitter in the central and peripheral nervous systems. Receptors for GABA have traditionally been divided into GABA<sub>A</sub> and GABA<sub>B</sub> receptor subtypes. GABA<sub>B</sub> receptors (for a review see Kerr, D.I.B. and Ong, J. (1995) Pharmac. Ther. vol. 67, pp.187-246) belong to the superfamily of G-protein coupled receptors. GABA<sub>B</sub> receptor agonists are described as being of use in the treatment of CNS disorders, such as muscle relaxation in spinal spasticity, cardiovascular disorders, asthma, gut motility disorders such as irritable bowel syndrome and as prokinetic and anti-tussive agents. GABA<sub>B</sub> receptor agonists have also been disclosed as useful in the treatment of emesis (WO 96/11680).
- The cloning of the rat GABA<sub>B</sub> receptors GABA<sub>B</sub>R1a (SEQ ID NOS: 44 and 45) and GABA<sub>B</sub>R1b (SEQ ID NOS: 46 and 47) is disclosed by Kaupmann et al. (1997) Nature, vol. 386, 239-246. The mature rat GABA<sub>B</sub>R1b differed from GABA<sub>B</sub>R1a in that the N-terminal 147 residues were replaced by 18 different residues. It was presumed that the rat GABA<sub>B</sub>R1a and -b receptor variants are derived from the same gene by alternative splicing.

The cloning of the human GABA<sub>B</sub> receptor GABA<sub>B</sub>R1b is disclosed in WO 97/46675.

#### Reflux

- In some humans, the lower esophageal sphincter (LES) is prone to relaxing more frequently than in other humans. As a consequence, fluid from the stomach can pass into the esophagus since the mechanical barrier is temporarily lost at such times, an event hereinafter referred to as "reflux".
- Gastro-esophageal reflux disease (GERD) is the most prevalent upper gastrointestinal tract disease. Current therapy has aimed at reducing gastric acid secretion, or by reducing esophageal acid exposure by enhancing esophageal clearance, lower esophageal sphincter tone and gastric emptying. The major mechanism behind reflux has been considered to depend on a hypotonic lower esophageal sphincter. However, recent research (e.g.
   Holloway & Dent (1990) Gastroenterol. Clin. N. Amer. 19, 517-535) has shown that most reflux episodes occur during transient lower esophageal sphincter relaxations (TLESR), i.e. relaxations not triggered by swallows. It has also been shown that gastric acid secretion usually is normal in patients with GERD. Consequently, there is a need for compounds

which reduce the incidence of TLESR and thereby prevent reflux.

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#### DISCLOSURE OF THE INVENTION

In the applicants' earlier patent application WO 98/11885, filed on 15 September 1997, it is disclosed that GABA<sub>B</sub> receptor agonists can be used to reduce the incidence of transient lower esophageal sphincter relaxations (TLESR).

The present invention provides nucleic acid molecules encoding human and canine GABA<sub>B</sub> receptors. These nucleic acid molecules will make possible the screening for compounds which are agonists or antagonists of GABA<sub>B</sub> receptors, e.g. compounds which are inhibitors of transient lower esophageal sphincter relaxations (TLESR).

Consequently, in a first aspect, the present invention provides a nucleic acid molecule encoding a GABA<sub>B</sub> receptor, or a functionally equivalent modified form thereof, said receptor being selected from the group consisting of human and canine GABA<sub>B</sub> receptors.

- In preferred forms of the invention, the said nucleic acid molecule encodes the human GABA<sub>B</sub> receptor 1a (SEQ ID NOS: 48 and 49), 1b (SEQ ID NOS: 50 and 51), 1c (SEQ ID NOS: 54 and 55) or 1d (SEQ ID NOS: 56 and 57); or the canine GABA<sub>B</sub> receptor 1a (SEQ ID NOS: 52 and 53) or 1c (SEQ ID NOS: 58 and 59). Accordingly, the invention furthermore provides a nucleic acid molecule selected from:
- 10 (a) nucleic acid molecules comprising a nucleotide sequence set forth as SEQ ID NO: 48, 50, 52, 54, 56 or 58;
  - (b) nucleic acid molecules comprising a nucleotide sequence capable of hybridizing, under stringent hybridization conditions, to a nucleotide sequence complementary to the polypeptide coding region of a DNA molecule as defined in (a); and
- (c) nucleic acid molecules comprising a nucleotide sequence which is degenerate as a result of the genetic code to a nucleotide sequence as defined in (a) or (b).

Furthermore, the invention provides nucleic acid molecules of genomic origin encoding human GABA<sub>B</sub> receptors (SEQ ID NOS: 60 and 61) as well as nucleic acid molecules (set forth as SEQ ID NO: 70, 72, 74, 76, 78, 80, 82, 84) encoding additional isoforms of the human GABA<sub>B</sub> receptor which isoforms can be shown to be generated by alternative splicing.

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It should thus be understood that the nucleic acid molecule according to the invention is not to be limited strictly to molecules comprising the sequences set forth as SEQ ID: 48, 50, 52, 54, 56 or 58. Rather the invention encompasses nucleic acid molecules carrying modifications like substitutions, small deletions, insertions or inversions, which nevertheless encode proteins having substantially the biochemical activity of the GABAB receptors according to the invention. Included in the invention are consequently nucleic acid molecules, the nucleotide sequence of which is at least 95% homologous, preferably at least 96%, 97%, 98% or 99% homologous, with the nucleotide sequence shown as SEQ ID NO: 48, 50, 52, 54, 56 or 58 in the Sequence Listing.

The term "stringent hybridization conditions" is known in the art from standard protocols (e.g. Current Protocols in Molecular Biology, editors F. Ausubel et al., John Wiley and Sons, Inc. 1994) and could be understood as as stringent or more stringent than those defined by e.g. hybridization to filter-bound DNA in 0.5 M NaHPO<sub>4</sub>, 7% sodium dodecyl sulfate (SDS), 1 mM EDTA at +65°C, and washing in 0.1xSSC / 0.1% SDS at +68°C.

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Included in the invention is also a nucleic acid molecule which nucleotide sequence is degenerate, because of the genetic code, to a nucleic acid of the present invention and more particularly to one of the nucleotide sequences set forth as SEQ ID NOs: 48, 50, 52, 54, 56 and 58. A sequential grouping of three nucleotides, a "codon", codes for one amino acid. Since there are 64 possible codons, but only 20 natural amino acids, most amino acids are coded for by more than one codon. This natural "degeneracy", or "redundancy", of the genetic code is well known in the art. It will thus be appreciated that the nucleic acid sequences shown in the Sequence Listing is only an example within a large but definite group of nucleic acid sequences which will encode the polypeptide as described above.

In a further aspect, the invention provides a recombinant polypeptide encoded by a nucleotide sequence of the present invention, encoding a GABA<sub>B</sub> receptor. In preferred forms of the inventions, the said polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 49, 51, 53, 55, 57, 59, 71, 73, 75, 77, 79, 81, 83 or 85 in the Sequence Listing. However, polypeptides of the present invention are not to limited to those having an amino acid sequence identical with one of SEQ ID NOs: 49, 51, 53, 55, 59, 71, 73, 75, 77, 79, 81, 83 or 85 in the Sequence Listing. Rather the invention encompasses polypeptides carrying modifications such as substitutions, small deletions, insertions or inversions, which polypeptides nevertheless have substantially the biological activities of the GABA<sub>B</sub> receptor. Included in the invention are consequently polypeptides, the amino acid sequence of which is at least 95% homologous, preferably at least 96%, 97%, 98% or 99% homologous, with one of the amino acid sequences described by SEQ ID NOs: 49, 51, 53, 55, 57 59, 71, 73, 75, 77, 79, 81, 83 and 85 in the Sequence Listing.

Included in the invention are polypeptides of the present invention which have been post-translationally modified, e.g. by cleavage of an N-terminal signal sequence which can be e.g. 1 to 25 amino acids long.

In yet another aspect, the invention provides a vector transformed with a nucleic acid molecule of the present invention. The said vector can e.g. be a replicable expression vector which carries and is capable of mediating the expression of a nucleic acid molecule according to the invention. In the present context the term "replicable" means that the vector is able to replicate in a given type of host cell into which is has been introduced.

Examples of vectors are viruses such as bacteriophages, cosmids, plasmids and other recombination vectors. Nucleic acid molecules are inserted into vector genomes by

methods well known in the art.

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Included in the invention is also a cultured host cell harbouring a vector according to the
invention. Such a host cell can be a prokaryotic cell, a unicellular eukaryotic cell or a cell
derived from a multicellular organism. The host cell can thus e.g. be a bacterial cell such as
an E. coli cell; a cell from a yeast such as Saccharomyces cervisiae or Pichia pastoris, or a
mammalian cell. The methods employed to effect introduction of the vector into the host
cell are standard methods well-known to a person familiar with recombinant DNA
methods.

A further aspect of the invention is a process for production of a GABA<sub>B</sub> receptor polypeptide according to the invention, said process comprising culturing a host cell as defined above under conditions whereby the said polypeptide is produced, and recovering the said polypeptide.

A further important aspect of the invention is a method for the screening of compounds which are inhibitors of transient lower esophageal sphincter relaxations (TLESR), said method comprising the use of a nucleic acid molecule encoding a GABA<sub>B</sub> receptor. The said nucleic acid molecule encoding a GABA<sub>B</sub> receptor can e.g. be one of the nucleic acid molecules according to the invention encoding human or canine GABA<sub>B</sub> receptors. However, it should be understood that this aspect of the invention is not limited to the use

of the said human and canine GABA<sub>B</sub> receptors, but rather encompasses the use of any GABA<sub>B</sub> receptor for screening for compounds which are inhibitors of transient lower esophageal sphincter relaxations.

In yet another important aspect, the invention provides a method for the screening of compounds which are agonists or antagonists to a GABA<sub>B</sub> receptor, said method comprising the use of a nucleic acid molecule, according to the invention, encoding human or canine GABA<sub>B</sub> receptors.

#### 10 Brief Description of the Drawings

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Figure 1: Map of the human GABA<sub>B</sub> receptor gene.

The exon/intron organisation is shown. Exons are indicated as solid boxes numbered 1-23.

The part of intron 5 that is retained together with exon 6 giving rise to GABA<sub>B</sub> receptor 1b is indicated as an open box.

Figure 2: Expression of human GABA<sub>B</sub> receptor 1b isoform in transfected C127 cells.

Western blot analysis of transfected C127 cells using a polyclonal anti-human GABA<sub>B</sub> receptor antibody. Lane 1: Untransfected C127 whole cell lysate. Lanes 2-7: Whole cell lysates of six independent clones transfected with human GABA<sub>B</sub> receptor 1b isoform encoding cDNA. The clones analysed in lanes 4 to 7 express a GABA<sub>B</sub> receptor of expected molecular weight (arrow).

Figure 3: Expression of human GABA<sub>B</sub> receptor 1d isoform in transfected C127 cells.

Western blot analysis of transfected C127 cells using a polyclonal anti-human GABA<sub>B</sub> receptor antibody. Lanes 1-3: Concentrated culture media from three independent C127 clones transfected with a cDNA expression construct encoding the human GABA<sub>B</sub> receptor

1d isoform. Lanes 4-6: Whole cell lysates corresponding to the clones analysed in lanes 1-3. The experiment revealed that the human GABA<sub>B</sub> receptor 1d cDNA encodes a secreted isoform. The arrow indicates the bands corresponding to the 1d isoform.

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Figure 4: Expression of human GABA<sub>B</sub> receptor 1d isoform in E. coli.

Western blot analysis of transformed *E. coli* cells using a polyclonal anti-human GABA<sub>B</sub> receptor antibody. Lane 1: Lysate from an uninduced *E. coli* culture transformed with an pET-based expression construct encoding the human GABA<sub>B</sub> receptor 1d cDNA. Lane 2: Lysate from an IPTG-induced *E. coli* culture transformed with an expression construct encoding the human GABA<sub>B</sub> receptor 1d cDNA. Lane 3: Lysate from an IPTG-induced *E. coli* culture transformed with an expression construct encoding an unrelated protein. Lane 4: The BSA-conjugated peptide previously used for immunization was loaded on the gel as a positive antibody control.

The screening methods according to the invention can e.g. comprise the steps (a) transforming a cultured cell with a nucleic acid molecule encoding a GABA<sub>B</sub> receptor, so that a GABA<sub>B</sub> receptor is expressed on the surface of the cell; (b) contacting a test compound with the said cell; and (c) determining whether the test compound binds to, and/or activates, the GABA<sub>B</sub> receptor.

In particular GABA<sub>B</sub> receptor expressing cells, transgenic animals or cells and tissues derived thereof, nay be used to screen substance libraries for antagonist or agonist activities. For this purpose, GABA<sub>B</sub> receptor expression may be directed to cells and tissues containing, either naturally or artificially, the necessary components allowing correct receptor transport and processing as well as coupling to second messenger pathways. Screening may be performed as ligand binding assays or functional assays. For screening, cells and tissues my be prepared in various ways, each uniquely suited to its purpose. Ligand binding assays are performed *in vivo* or *in vitro* using e.g. radiolabelled GABA. Functional assays examplified by, but not limited to, Ca<sup>++</sup>-responses, cAMP-

responses and effects on K<sup>+</sup> channels, may be performed in living cells, broken cells, isolated cell membranes, as well as in tissues and in living animals. To facilitate measurement of physiological GABA<sub>B</sub> receptor mediated responses, GABA<sub>B</sub> receptors may be co-expressed with promiscous G-proteins like e.g. Ga16 or Gqi5 increasing G-protein coupling. Another possible way to increase G-protein coupling is to fuse the GABA<sub>B</sub> receptor with appropriate G-proteins using standard molecular techniques. In order to further improve readouts in Ca<sup>++</sup>-response assays, GABA<sub>B</sub> receptors may be co-expressed with aequorin, a photoprotein cloned from the luminescent jellyfish Aequorea

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In yet another aspect, the invention provides a pharmaceutical compositions comprising a soluble GABA<sub>B</sub> receptor further comprising one or more pharmaceutical acceptable carriers and/or diluents.

In yet another aspect, the invention provides a method for the diagnosis or treatment of conditions related to GABA-dysfunction, e.g. epilepsy, psychiatric disorders such as depression and anxiety, cognitive dysfunction, gastroesophageal reflux disease, emesis, irritable bowel syndrome, dyspepsia, spasticity, arthritis, allergies, autoimmune diseases, neoplastic diseases, pain, and infectious diseases, comprising the use or administration of soluble forms of the GABA<sub>B</sub> receptor, such as the human GABA<sub>B</sub> receptor 1c or 1d or functionally equivalent modified forms thereof; preferably being a polypeptidfe of the present invention.

The soluble forms of the GABA<sub>B</sub> receptor can e.g. be produced by culturing a host cell harbouring a vector comprising a nucleic acid encoding the soluble GABA<sub>B</sub> receptor under conditions where whereby the said polypeptide is produced, the polypeptide recovered and administered to a patient in need thereof.

Furthermore, the level of soluble GABA<sub>B</sub> receptors in certain body fluids, e.g. serum and CSF, can be measured and used in diagnostics related to conditions with altered levels of soluble GABA<sub>B</sub> receptors in said body fluids.

Throughout this description the terms "standard protocols" and "standard procedures", when used in the context of molecular cloning techniques, are to be understood as protocols and procedures found in an ordinary laboratory manual such as: Current Protocols in Molecular Biology, editors F. Ausubel et al., John Wiley and Sons, Inc. 1994, or Sambrook, J., Fritsch, E.F. and Maniatis, T., Molecular Cloning: A laboratory manual, 2nd Ed., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY 1989.

#### **EXAMPLES**

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EXAMPLE 1A: Cloning and sequencing of cDNA encoding human GABA<sub>B</sub> receptor 1a and 1b

Messenger RNA from human hippocampus was obtained from Clontech (Palo Alto, CA, USA) (catalog #6578-1). First-strand cDNA synthesis reactions were performed using the First-strand cDNA Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden). The pd(N)<sub>6</sub> primer was used to prime the first-strand synthesis. The generated cDNA molecules were used as template in the PCR reactions described below.

Specific PCR primers were designed (Table 1) based on the sequences of the rat GABAB receptor 1a and 1b cDNA (Kaupmann et al. 1997, EMBL accession numbers Y10369 (SEQ ID NO: 44) and Y10370 (SEQ ID NO: 46)).

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TABLE 1
Primers used for RT-PCR on mRNA from human hippocampus

Nr.	Species	Sequence 5'- 3'	SEQ ID NO	
794	Rat	GTTTCTTCTCGGATCCAGCTGTGCCTG	1	
795	Rat	CAGGCACAGCTGGATCCGAGAAGAAACT	2	
796	Rat	CGGTCGACTCACTTGTAAAGCAAATGTACTCGACTCCC	3	
797	Rat	ATGCGCGCCGGCAGCCAACATGCTGCTGCTGCTGCTGCTGC	4	
831	Rat	CGGTCGACTCACTTGTAAAGCAAATGTACTCGACTCCCATCACAGC	5	
838	Rat	ATGCGCGCCGGCAGCCAACATGCTGCTGCTGCTGCTGCTGCTCTCTTCC	6	
842	Rat	CAGGCACAGCTGGATCCGAGAAGAAACTCTGTCGGAAAGT	7	
863	Rat	GGTCATCCAGCGTTGAGGTGAAGAC	8	
864	Rat	GAAGGTTGCCAGATTATACATCCGC	9	
865	Rat	CCACGATGATTCGAGCATCTTGACG	10	
866	Rat	GCCTCTCACTCCCCTCATCTCC	11	
932	Human	GAGTGAAGGAGGCTGGAATTG	12	

cDNA fragments encoding parts of the human GABA<sub>B</sub> receptors were amplified directly by PCR using the designed primers with the generated cDNA molecules as templates. All PCR experiments were carried out using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular Systems Inc., NJ, USA) with the following PCR program: +95°C for 1 minute, +50°C for 30 seconds, +72°C for 3 minutes, repeated 44 times and finally +72°C for 7 minutes. The following primer combinations gave PCR products with expected sizes: 838 and 842, 838 and 795, 797 and 865, 864 and 865, 864 and 863 which corresponds to the 5′-end of the GABA<sub>B</sub> receptor 1a. Primer pair 932 and 831, 932 and 796, 794 and 831 gave PCR products which correspond to the 3′-end of both GABA<sub>B</sub> receptor 1a and 1b. The primer combination 839 and 918 gave a PCR product
 corresponding to the 5′-end of the GABA<sub>B</sub> receptor 1b.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to the cDNA encoding the GABA<sub>B</sub> receptor were used.

Additional PCR primers were designed based on the obtained sequences encoding fragments of the human GABA<sub>B</sub> receptor and additional DNA fragments encoding parts of the human GABA<sub>B</sub> receptors were amplified by PCR, the PCR products subcloned and sequenced as described above.

EXAMPLE 1B: Cloning and sequencing of the 3'-ends of the cDNA encoding human GABA<sub>B</sub> receptors 1a and 1b

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Messenger RNA from human hippocampus was obtained from Clontech (Palo Alto, CA, USA) (catalogue #6578-1). First strand cDNA synthesis reactions were performed using the First-strand cDNA Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden). The Not I-d(T)<sub>18</sub> primer was used to prime the first-strand synthesis. The generated cDNA molecules were used as template in the PCR reactions described below.

Specific PCR primers were designed (Table 2) based on the sequences of the human GABA<sub>B</sub> receptor 1a and 1b cDNA obtained in Example 1A and the EST sequence EMBL accession number Y11044.

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By homology searches in the EMBL database using the GABA<sub>B</sub> receptor cDNA sequences obtained in Example 1A as query the EST sequence EMBL accession number Y11044 have been found to be homologous to the 3'-end of the GABA<sub>B</sub> receptor cDNA.

## TABLE 2Primers used in PCR to amplify 3'ends of human GABA<sub>B</sub> receptor cDNA

Nr	Species	Sequence 5'-3'	SEQ ID NO
938	Human	GACGCTTATCGAGCAGCTTC	13
972	Human	AGCCCAGAACTCACAGGGGGACAT	14
973	Human	GCTTCAAGCCAGGTACGAACTAA	15

cDNA fragments encoding parts of the human GABA<sub>B</sub> receptors were amplified directly by PCR using the designed primers with the generated cDNA molecules as templates. All

PCR experiments were carried out using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular Systems Inc., NJ, USA) with the following PCR program: +95°C for 1 minute, +50°C for 30 seconds,+72°C for 3 minutes, repeated 44 times and finally +72°C for 7 minutes. The following primer combinations gave PCR products with expected sizes: 938 and 972, 938 and 973 corresponding to the 3'end of both GABAB receptor 1a and 1b cDNA.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to the cDNA encoding the GABA<sub>B</sub> receptor were used.

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EXAMPLE 1C: Cloning and sequencing of the 5'-end of the cDNA encoding human GABA<sub>B</sub> receptor 1b

20 Messenger RNA from human hippocampus was obtained from Clontech (Palo Alto, CA, USA) (catalogue #6578-1). Marathon cDNA amplification Kit (Clontech) was used for performing 5'/3'- RACE (Rapid Amplification of cDNA Ends). Adaptor-ligated double stranded cDNA molecules were amplified according to the manufacturers description. The pd(N)<sub>6</sub> primer from the First-strand cDNA Synthesis kit from Amersham Pharmacia
25 Biotech (Uppsala, Sweden) was used to produce the adaptor-ligated cDNA.

A specific PCR primer was designed (Table 3) based on the sequences of the human GABA<sub>B</sub> receptor 1b cDNA obtained in Example 1A.

TABLE 3
Primers used in PCR to amplify the 5'-ends of human GABA<sub>B</sub> receptor cDNA

Nr.	Species	Sequence 5'-3'	SEQ ID NO
958	Human	TGGCCCTCACCGCCTCAGTCATCTCA	16
API	Marathon kit	CCATCCTAATACGACTCACTATAGGGC	17

5 cDNA fragments encoding part of the human GABA<sub>B</sub> receptors were amplified directly by PCR using the designed primers with the generated adaptor-ligated cDNA molecules as template. The PCR experiment was carried out using the Expand Long Template PCR System (Boehringer Mannheim GmbH, Germany) with the following PCR program: + 94°C for 1minute, + 94°C for 30 seconds, +60°C for 30 seconds, +68°C for 4 minutes, repeated 24 times. The primer combination AP1 and 958 gave a PCR product which corresponded to the 5′-end of the GABA<sub>B</sub> receptor 1b cDNA, including 190 base pairs upstream the initiation codon.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison,
USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide
sequence for all subclones were determined using a Thermo Sequenase dye terminator
cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As
primers for sequencing reactions specific oligonucleotides complementary to the vector
pGEM-T or primers complementary to the cDNA encoding the GABAB receptor were
used.

EXAMPLE 1D: Cloning and sequencing of the 5'-end of the cDNA encoding human GABA<sub>B</sub> receptor 1a

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Messenger RNA from human hippocampus was obtained from Clontech (Palo Alto, USA) (catalogue #6578-1). Marathon cDNA amplification Kit (Clontech) was used to obtain adaptor-ligated double stranded cDNA molecules according to the manufacturer's description. The pd(N)<sub>6</sub> primer from the First-strand cDNA Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden) was used to obtain the adaptor-ligated cDNA.

Specific PCR primers were designed (Table 4) based on the sequences of the human GABA<sub>B</sub> receptor 1a cDNA obtained in Example 1 and the rat GABA<sub>B</sub> receptor 1a cDNA disclosed in WO 97/46675.

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TABLE 4
Primers used to amplify 5'-ends of the human GABAB receptor 1a cDNA

Nr.	Species	Sequence 5'-3'	SEQ ID NO
1033	Human	CTCAATCTCATAGTCCACTGG	18
1087	Rat	CCTTGAGGCCCGGGGAGAG	19

cDNA fragments encoding part of the human GABAB 1a receptor were amplified directly by PCR using the designed primers with the generated adaptor-ligated cDNA molecules as template. The PCR was performed using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular Systems Inc., NJ, USA) with the following PCR program: +94°C for 1 minute, +50°C for 30 seconds,+72°C for 3 minutes; +94°C for 1 minute, +60°C
 for 30 seconds,+72°C for 4 minutes, repeated 34 times and finally +72°C for 7 minutes.

The primer combination 1087 and 1033 gave a PCR product corresponding to the 5'-end of the GABA<sub>B</sub> receptor 1a cDNA, including 26 base pairs upstream the initiation codon.

20 The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T were used.

A complete cDNA sequence encoding the human GABA<sub>B</sub> receptors 1a (SEQ ID NO: 48) and the human GABA<sub>B</sub> receptor 1b (SEQ ID NO: 50) were obtained by aligning the sequences of the different fragments cloned and sequenced in Examples 1A, 1B, 1C and 1D.

EXAMPLE 2A: Cloning and sequencing of cDNA encoding canine GABA<sub>B</sub> receptor 1a

- QuickPrep Micro mRNA Purification kit (Amersham Pharmacia Biotech, Uppsala, Sweden) was used to isolate mRNA from canine neural tissues according to the manufacturers description. First-strand cDNA synthesis reactions were performed using the First-strand cDNA Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden). The Not-I-d(T)<sub>18</sub> bifunctional or pd(N)<sub>6</sub> primer was used to prime the first-strand synthesis.
  The generated cDNA molecules were used as template in the PCR reactions described
- below.

Specific PCR primers (Table 5) were designed based on the sequences of the rat GABA<sub>B</sub> receptor 1a and 1b cDNA (Kaupmann et al. 1997, EMBL accession numbers Y10369 (SEQ ID NO: 44 and Y10370 (SEQ ID NO: 46)).

TABLE 5
Primers used for RT-PCR on mRNA from canine cortex

Nr.	Species	Sequence 5'-3'	SEQ ID NO
795	Rat	CAGGCACAGCTGGATCCGAGAAGAAACT	20
831	Rat	CGGTCGACTCACTTGTAAAGCAAATGTACTCGACTCCCATCACAGC	21
838	Rat	ATGCGCGCCGGCAGCCAACATGCTGCTGCTGCTGCTGCTCTCTTCC	22
840	Rat	CGTCAAGATGCTCGAATCATCG	23
841	Rat	CAGGGGCTCAGAGGTCCC	24
842	Rat	CAGGCACAGCTGGATCCGAGAAGAACTCTGTCGGAAAGT	25
844	Rat	CGGTCGACTCACTTGTAAAGCAAATGTACTCGACTCCCATCACAGCTAAG	26
848	Rat	ACTTTCCGACAGAGTTTCTTCTCGGATCCAGCTGTGCCTG	27
865	Rat	CCACGATGATTCGAGCATCTTGACG	28

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cDNA fragments encoding parts of the canine GABA<sub>B</sub> receptor were amplified directly by PCR using the designed primers with the generated cDNA molecules as templates. All PCR experiments were carried out using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular Systems Inc., NJ, USA) with the following PCR program: +95°C for 1 minute, +50°C for 30 seconds, +72°C for 3 minutes, repeated 44 times and finally

+72°C for 7 minutes. The following primer combinations gave PCR products with expected sizes: 842 and 838, 838 and 795, 838 and 865 which corresponds to the 5'-part of the canine GABA<sub>B</sub> receptor cDNA. Primer pair 848 and 844, 848 and 831, 848 and 841, 840 and 841 gave PCR products which correspond to the 3'-part of the canine GABA<sub>B</sub> receptor cDNA.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to the cDNA encoding the GABA<sub>B</sub> receptor were used.

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EXAMPLE 2B: Cloning and sequencing of the 3'- and 5'-ends of the cDNA encoding canine GABA<sub>B</sub> receptor 1a

QuickPrep Micro mRNA Purification kit (Amersham Pharmacia Biotech, Uppsala,

Sweden) was used to isolate mRNA from canine nerve tissues according to the
manufactures description. Marathon cDNA amplification Kit (Clontech, Palo Alto, CA,
USA) were used for performing both 5'-and 3'-RACE. Two adaptor-ligated double
stranded cDNA libraries were amplified according to the manufacturers description. A
random primer (pd(N)6) was used when amplifying the adaptor-ligated cDNA for the 5'
RACE and the Marathon cDNA Synthesis primer (52-mer) was used when amplifying the

adaptor ligated cDNA for the 3'-RACE.

Specific PCR primers were designed (Table 6) based on the sequence of canine GABA<sub>B</sub> receptor 1a cDNA obtained in Example 2A.

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TABLE 6
Primers used in PCR to amplify the 5'- and 3'-ends of canine GABA<sub>B</sub> receptor 1a cDNA

Nr.	Species	Sequence 5'-3'	SEQ ID NO
936	canine	CTACCGCGCAATGAACTCCTCGTC	29
1076	canine	CGAGGTGGCGTTGGGGGTCTGTGC	30
AP1	Marathon kit	CCATCCTAATACGACTCACTATAGGGC	31
AP2	Marathon kit	ACTCACTATAGGGCTCGAGCGGC	32

5 cDNA fragments encoding parts of the canine GABA<sub>B</sub> receptor were amplified by PCR from the adaptor-ligated cDNA using the designed primers. A number of different PCR programs were tested to find conditions where PCR products corresponding to GABA<sub>B</sub> receptor DNA were obtained. The 5'- PCR experiments were carried out using the Expand Long Template PCR System (Boehringer Mannheim GmbH, Germany) with the following PCR program: +94°C for 30 seconds, +72°C for 3 minutes, repeated 4 times; +94°C for 30 seconds, +68°C for 3 minutes, repeated 24 times. The primer combination AP2 and 1076 gave a PCR product which corresponded to the 5'-end of the GABA<sub>B</sub> receptor cDNA, including 114 base pairs upstream the initiation codon.

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The 3'- PCR experiments were carried out using the Expand Long Template PCR System (Boehringer Mannheim GmbH, Germany) with the following PCR program: +94°C for 1 minute; and +94°C for 30 seconds, +60°C for 30 seconds, +68 °C for 4 minutes, repeated 29 times. The primer combination AP1 and 936 gave a PCR fragment which corresponded to the 3'-end of the GABA<sub>B</sub> receptor cDNA, including the poly(A) tail.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit(Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to GABAB receptor DNA were used.

Complete cDNA sequence encoding the canine GABA<sub>B</sub> receptor 1a (SEQ ID NO: 52) was obtained by aligning the sequences of the different fragments obtained in Example 2A and Example 2B.

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EXAMPLE 3A: Cloning of cDNA encoding human GABA<sub>B</sub> receptor 1c and 1d from Jurkat cells

A guanidinisothiocyanate/CsCl method was used to isolate total RNA from Jurkat cells.

The first-strand cDNA synthesis was performed using the First-strand cDNA Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden). The pd(N) primer was used to prime the first strand synthesis. The generated cDNA molecules were used as templates in the PCR reaction described below.

Specific PCR primers (Table 7) were designed based on the sequence of human GABA<sub>B</sub> receptor 1a and 1b cDNA (Example 1), rat GABA<sub>B</sub> receptor (Kaupmann et al. 1997) and the EST sequence EMBL accession number Y11044.

TABLE 7

20 Primers used in RT-PCR on mRNA from Jurkat cells

Nr	Species	Sequence 5'-3'	SEQ ID NO
938	human	GACGCTTATCGAGCAGCTTC	33
972	human	AGCCCAGAACTCACAGGGGGACAT	34
973	human	GCTTCAAGCCAGGTACGAACTAA	35
893	rat	GGAGCACCCCAAGCCCCACTG	36
937	human	CTGGTTCCTCCCAATGTG	37
1005	rat	CCTCTCACTCCCCTCATCTC	38
1030	human	AAGCCAACCTTCCCTGCTTCTC	39

cDNA fragments encoding parts of the GABA<sub>B</sub> receptor were amplified directly by PCR using human and rat specific primers. All PCR experiments were carried out using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular Systems Inc., NJ,

USA) with the following PCR program: +95°C for 1 minute; +54°C for 1 minutes, +72°C for 3 minutes, repeated 44 times; and finally +72°C for 7 minutes.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to GABA<sub>B</sub> receptor DNA were used.

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The following primer combination gave PCR products corresponding to the 3'-end of the GABA<sub>B</sub> receptor cDNA: primer pairs 938 and 972; 938 and 973. Unexpectedly both these fragments lacked 149 base pairs, resulting in a frame shift and the insertion of a new termination codon. The following primer combination gave a PCR product corresponding to the 5'-part of the GABA<sub>B</sub> receptor 1a cDNA: 893 and 937. The primer pairs 1005 and 937, 1030 and 937 gave PCR products corresponding to the 5'-part of the GABA<sub>B</sub> receptor 1b cDNA. Also these PCR fragments lacked the same 149 base pairs which resulted in a frame shift and the insertion of a new termination codon.

These results show that Jurkat cells contain mRNA encoding two new forms of the human GABA<sub>B</sub> receptor. These are designated GABA<sub>B</sub> receptor 1c (SEQ ID NO: 54 and 55) (comprising the same 5'-part as the GABA<sub>B</sub> receptor 1a) and GABA<sub>B</sub> receptor 1d (SEQ ID NO: 56 and 57) (comprising the same 5'-part as the GABA<sub>B</sub> receptor 1b). These two forms of the GABA<sub>B</sub> receptor do not contain any of the transmembrane regions and are therefore expected to be soluble forms of the receptor.

EXAMPLE 3B: Analysis of cDNA encoding human GABA<sub>B</sub> receptors from hippocampus

Messenger RNA from human hippocampus was obtained from Clontech (Palo Alto, USA) (catalogue #6578-1). First strand cDNA synthesis reactions were performed using the First-strand cDNA Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden). The

pd(N)<sub>6</sub> primer was used to prime the first-strand synthesis. The generated cDNA molecules were used as template in the PCR reactions described below.

Specific PCR primer was designed (Table 8) based on the sequences of the cDNAs encoding human GABA<sub>B</sub> receptors 1a and 1b.

TABLE 8
Primers used for RT-PCR on mRNA from human hippocampus

Nr	Species	Sequence 5' - 3'	SEQ ID NO
937	Human	CTGGTTCCTCCCAATGTG	40
938	Human	GACGCTTATCGAGCAGCTTC	41

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cDNA fragments encoding parts of the human GABA<sub>B</sub> receptors were amplified directly by PCR using the designed primers with the generated cDNA molecules as templates. All PCR experiments were carried out using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular Systems Inc., NJ, USA) with the following PCR program: +94°C for 1 minute, +50°C for 30 seconds,+72°C for 3 minutes; +94°C for 1 minute, +54°C for 30 seconds, +72°C for 3 minutes repeated 44 times; and finally +72°C for 7 minutes. The primer combination 938 and 937 gave a PCR product which corresponded to the expected size of the GABA<sub>B</sub> receptor 1a and 1b cDNA but also a fragment a smaller size.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to the cDNA encoding the GABA<sub>B</sub> receptor were used.

The larger PCR fragment was found to correspond to the 3'-part of the GABA<sub>B</sub> receptor 1a and 1b cDNA and the smaller fragment which lacked 149 base pairs was found to correspond to the 3'-part of the GABA<sub>B</sub> receptors 1c and 1d cDNA identified in Example 3A.

EXAMPLE 4: Cloning and sequencing of cDNA encoding canine GABA<sub>B</sub> receptor 1b

cDNA encoding the canine GABA<sub>B</sub> receptor 1b is isolated in a similar manner as described in Example 2 for receptor 1a. PCR primers specifically designed to be complementary to the 5'-end of the cDNA encoding the rat and human GABA<sub>B</sub> receptor 1b, together with PCR primers complementary to the 3'-end of the cDNA encoding the canine GABA<sub>B</sub> receptor 1a, and mRNA prepared from a suitable canine tissue, are used.

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EXAMPLE 5: Cloning of cDNA encoding canine GABA<sub>B</sub> receptor 1c

Total RNA from canine liver was prepared using the RNeasy Total RNA Purification

Protocols (Quiagen GmbH, Germany). The first-strand cDNA synthesis was performed using the First-strand cDNA Synthesis kit from (Amersham Pharmacia Biotech, Uppsala, Sweden). The pd(N)<sub>6</sub> primer was used to prime the first-strand synthesis. The generated cDNA molecules were used as templates in the PCR reaction described below.

20 Specific PCR primers (Table 9) were designed based on the sequence of canine GABA<sub>B</sub> receptor 1a cDNA.

TABLE 9
Primers used in RT-PCR

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Nr	Species	Sequence 5'-3'	SEQ ID NO
936	canine	CTACCGCGCAATGAACTCCTCGTC	42
954	canine	CCTTCTTCTCCTCCTTCTTAGTGA	43

cDNA fragments encoding parts of the canine GABA<sub>B</sub> receptor were amplified directly by PCR using canine specific primers. All PCR experiments were carried out using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular Systems Inc., NJ, USA) with the following PCR program: +95°C for 1 minute, +54°C for 30 seconds, +72°C for 3

minutes, repeated 44 times and finally +72°C for 7 minutes. The primer combination gave a PCR product with a size corresponding to the GABA<sub>B</sub> receptor 1a but also a fragment of smaller size indicating the presence of a GABA<sub>B</sub> receptor 1c.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, USA). The inserts were subjected to nucleotide sequence analysis, the complete nucleotide sequence for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T were used.

The smaller fragment was shown to lack 149 base pairs. This deletion caused a frame shift and an insertion of a new termination codon, verifying the existence of a canine GABA<sub>B</sub> receptor 1c.

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Complete cDNA sequence encoding the canine GABA<sub>B</sub> receptor 1c (SEQ ID NO: 58) was obtained by aligning the sequences of the fragments obtained in Example 2A, Example 2B and Example 5.

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EXAMPLE 6. Cloning, sequencing and organization of human GABA<sub>B</sub> receptor genomic fragments

To determine the structural organization and sequence of the human GABA<sub>B</sub> receptor gene,

human genomic DNA libraries and human genomic DNA were screened and analyzed.

Human genomic libraries were obtained from Clontech (Palo Alto, CA, USA). The

libraries were constructed from female leukocyte DNA (catalog # HL1111J), cloned into

λEMBL-3 vector. The average size of inserts are 16 kb and the number of independent

clones are 1.7x10<sup>6</sup>. Human genomic DNA was obtained from Clontech (catalog # 6550-1).

In order to isolate recombinant phages containing exon and intron sequences of the human

GABA<sub>B</sub> receptor gene, 48 individual bacterial plates with a diameter of 150 mm and

approximately 4 x 10<sup>4</sup> individual plaques per plate, were screened. The methods and

solutions used were as described in the Library Protocol Handbook: General Procedures for the Hybridization of Lambda Phage Libraries w/DNA Probes (Clontech) with some modifications as will be apparent from the following.

5 The experiment was carried out essentially as follows. The numbers are given per plate basis. A sample of the phage library diluted in 0.1 ml sterile lambda diluent was prepared in order to obtain an estimated titer of 40,000 pfu (plaque forming units). A 0.6 ml LB-medium culture of the *E. coli* host strain K802 (obtained from Clontech) was infected with 40000 pfu recombinant phages for 15 minutes at +37°C. The culture was then mixed with 7 ml top agarose (6.5 g of agarose added per liter LB) and poured onto LB plates. The plates were incubated at +37°C for approximately 7 hours. The plates were then chilled at +4°C.

Plaque hybridization experiments were as follows. Membrane filters, Colony/Plaque Screen (DuPont, Wilmington, DE, USA), were placed onto the top of the plates for 3 minutes. For denaturation of DNA the filters were removed and floated in 0.5 M NaOH on a plastic wrap for 2 minutes, with the plaque side up. This step was repeated once to ensure efficient denaturation. Following neutralization the membrane filters were placed in 1M Tris-HCl pH 7.5, two times 2 minutes and allowed to dry.

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To obtain probes for DNA hybridization screening of the membrane filters, a GABA<sub>B</sub> receptor cDNA clone was digested with SacII and a 479 bp fragment (base pairs 573-1051 in the cDNA encoding human GABA<sub>B</sub> receptor 1a, SEQ ID NO: 48) was separated by agarose electrophoresis, excised and transferred to a polypropylene microcentrifuge tube. Additional probes were obtained by PCR amplification of various regions of the GABA<sub>B</sub> receptor cDNA (base pairs 68-486 and 2368-2863 in the cDNA encoding human GABA<sub>B</sub> receptor 1a, SEQ ID NO: 48). The isolated cDNA fragment was <sup>32</sup>P-labeled using Megaprime DNA labeling system (Amersham Pharmacia Biotech, Uppsala, Sweden) by the following procedure. Water was added at a ratio of 3 ml per gram of gel, and placed in a boiling water bath for 7 minutes to melt the gel and denature the DNA. A volume of DNA/agarose solution containing 25 ng of DNA was added to the labeling reaction, according to the supplier's instructions. Labeled nucleotides were removed from DNA

labeling reactions using MicroSpin<sup>TM</sup> G-50 Columns (Amersham Pharmacia Biotech, Uppsala. Sweden ).

The DNA hybridization reaction was performed under stringent conditions according to the method described below. The filter membranes were prehybridized at +65°C for at least 1 hour in a solution composed of 1% SDS, 1M NaCl, and 10% dextran sulfate using a hybridization oven (Hybaid Ltd, Ashford, UK). Following prehybridization a solution containing denatured herring sperm DNA of a final concentration of 100 µg/ml and the 32P-labeled DNA probe at a concentration <10 ng/ml (for optimal signal to background ratio) was added to the prehybridization solution and the membrane filters were incubated at +65°C for 10-20 hours. Following the removal of the hybridization solution the membrane filters were first washed in a 2xSSC (0.3M NaCl, 0.03M Na-citrate), 1% SDS solution two times for 5 minutes at room temperature. In the next step, the membrane filters were incubated +60°C two times for 30 minutes each in the same solution. In a third step, the filters were washed two times at room temperature in 0.1xSSC. Finally, the membrane filters were placed on a sheet of filter paper with the DNA face up, and allowed to dry. The dried membrane filters were then exposed to X-ray films and autoradiographed.

Of the approximately 2x10<sup>6</sup> individual plaques analyzed, four hybridizing plaques were detected and isolated. These three isolates were designated #GR1, #GR12, #GR13 and #GR41, respectively. After several rescreening experiments, the recombinant phage DNA was purified using Qiagen Lambda Midi Kit (Qiagen GmbH, Germany). The purified DNA was digested with SalI and the fragments representing the inserts were isolated by agarose electrophoresis.

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The sizes of the inserts were approximately isolate #GR1, 12 kb, isolate #GR12, 12 kb; isolate #GR13, 16kb; and isolate #GR41, 19 kb. These fragments were cloned into SalI digested linearized pUC19, resulting in the plasmids pAM362 (isolate #GR1), pAM363 (isolate #GR12), pAM364 (isolate #GR13)and pAM365 (isolate #GR41). The inserts from the four plaques which hybridizes to the GABA<sub>B</sub> receptor cDNA probes were analyzed by PCR, restriction mapping and hybridization to <sup>32</sup>P-labeled DNA fragments representing various regions of the GABA<sub>B</sub> receptor gene.

The cloned fragments in the plasmids pAM362, pAM363, pAM364 and pAM365 were characterized by restriction enzyme mapping, using EcoRI, HindIII, PstI, and BamHI. The approximate positions of the exons and the approximate size of the introns were analyzed and determined by PCR-based exon-exon linking and agarose gel electrophoresis.

In order to facilitate nucleotide sequence analysis, 7 restriction sub-fragments derived from pAM364, 2 restriction fragments derived from pAM362, and 1 restriction sub-fragment derived from pAM365 were isolated and cloned into pUC19, resulting in the plasmids pAM366-pAM375. The following strategy was employed; by combining PCR primers located within the pUC19 sequence either upstream or downstream of the cloning site, with a PCR primer with defined orientation and specific for the GABA<sub>B</sub> receptor derived subcloned fragment allowed the determination.

The inserts in the 10 plasmids pAM366-pAM375 were subjected to nucleotide sequence analysis. The nucleotide sequences for all subclones were determined using a Thermo Sequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As primers for sequencing reactions specific oligonucleotides complementary to pUC19 or primers complementary to the GABA<sub>B</sub> receptor cDNA were used.

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The genomic fragments cloned in the plasmids pAM362-pAM365 where shown to contain the complete transcribed part of the human GABA<sub>B</sub> receptor gene and extends > 3kb upstream of the first exon and > 2kb downstream of the last exon. The fragments cloned in the plasmid pAM362 were shown to contain exons 7-11, pAM363 exons 12-22, pAM364 exons 1-11, and pAM365 exons 12-23 of the GABA<sub>B</sub> receptor gene (Figure 1). The sequence of exons 1-11 and introns 1-10 is listed in SEQ ID NO: 60 and the sequence of exons 12-23 and introns 12-22 is listed in SEQ ID NO: 61.

The human GABA<sub>B</sub> receptor gene consists of 23 exons and 22 introns (Figure 1). The size of the exons range between 21 bp and 1486 bp. As can be noted from Table 10 exon/intron boundaries are in accordance with the AG/GT rule and conform well to the consensus sequence suggested by Mount et al. 1982.

TABLE 10

Exon-Intron boundaries of the GABA<sub>B</sub> receptor gene, sequences at exon-intron junctions.

		5'splice donor	3'splice acceptor	
Exon 1-Exon 2	CGAG	GTAAGAG	CCGCCTCTCACTTAG	ATGT
Exon 2-Exon 3	GAAG	GTGCATC	CGACTCACCCCTTAG	GTTG
Exon 3-Exon 4	TGTG	<del>_</del>	CCWATCTCTCCAC <u>AG</u>	TCCG
Exon 4-Exon 5	CAGG	GTGAGGG	CTTTCCTGCTGCCAG	TGAA
Exon 5-Exon 6	TCAG	GTGAGAT	CGCACCCCTCCTCAG	AACG
Exon 6-Exon 7	CAAG	GTAGCCC	CCTCTTGTCTTCAG	TGTG
Exon 7-Exon 8	TGTG		CTCCCTGCCCCACAG	CTTT
Exon 8-Exon 9	TTCG		TTATTCCCACCCAAG	ACTC
Exon 9-Exon10	GAAG	<del></del>	CTTTCTCTGTKGTAG	CGCC
Exon10-Exon11	TGAG		CTCCTCTGTATTCAG	GTGT
Exon11-Exon12	CATG	GTGAGAG	TTTTTTCCTCCCAAG	ACAT
Exon12-Exon13	CTCT	GTGAGTT	TGTTCCTTCCTCAG	GGCC
Exon13-Exon14	CAGG	<u>GT</u> TAGTA		GTGG
Exon14-Exon15	ATTG		CCCTGTGCCATGCAG	GAGG
Exon15-Exon16	TCCG		CCACCTCTGCCCTAG	TTAT
Exon16-Exon17	CCAG		TCTCTTCCTTTCTAG	GCCC
Exon17-Exon18	GAAG		CACATATTTATCCAG	ACTC
Exon18-Exon19	TGAG		TYGTTTCTGCCCTAG	ACAT
Exon19-Exon20	CTTG	_	CTCCTGCCATCCTAG	GCAT
Exon20-Exon21	GGCA	GTGAGCA	TGTCTTTCCCTCTAG	GTCC
Exon21-Exon22	CAAG		AACATTTGCCCCCAG	ATGC
Exon22-Exon23	TGAG	GTGCGGG	TGCTTCTTCCTCCAG	AAAG

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A comparison of the sequences of the different forms of the human GABA<sub>B</sub> receptor cDNA (SEQ ID NO: 48, 50, 54, and 56) and the sequence of the human GABA<sub>B</sub> gene (SEQ ID NO: 60 and 61) reveals that different mRNA encoding human GABA<sub>B</sub> receptors are formed by alternative splicing. The translational start of the GABA<sub>B</sub> receptor 1a is localized in exon 2 and the translational stop is localized in exon 23. The mRNA encoding GABA<sub>B</sub> receptor 1b is formed by alternative splicing where part of intron 5 is retained together with exon 6 where the translational start of the GABA<sub>B</sub> receptor 1b is derived from the intron sequence. The mRNA encoding GABA<sub>B</sub> receptor 1c is formed by alternative splicing where exon 15 are removed together with introns 14 and 15 and a frame shift and a translational stop are generated in the sequence corresponding to exon 16.

The mRNA encoding GABA<sub>B</sub> receptor 1d is formed when the translational start of the GABA<sub>B</sub> receptor 1b is generated together with the translational stop of the GABA<sub>B</sub> receptor 1c.

The la isoform is formed by splicing exon 5 to a cryptic splice site in the middle of exon 6.

Transcription of 1b isoform mRNA is most likely initiated from regulatory elements in intron 5. The ATG initiating translation of 1b mRNA is located in the 5'-end of exon 6.

Additional variants of the mRNA encoding variants of the human GABA<sub>B</sub> receptor can be derived by alternative splicing where one ore more of the exons or part of exons are excised in the processing of the pre-mRNA. The subsequent transcription of these mRNA will give rise to variants of the human GABA<sub>B</sub> receptor with potentially different biological and/or pharmacological activities.

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EXAMPLE 7. Analysis of cDNA encoding human GABA<sub>B</sub> receptors from human brain

Messenger RNA from human fetal brain (catalog #6525-1) and adult human brain (catalog #6516-1) were obtained from Clontech (Palo Alto, CA, USA). First strand cDNA synthesis reactions were performed using the first strand cDNA Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden). The pd(N)<sub>6</sub> primer was used to prime the first-strand synthesis. The generated cDNA molecules were used as templates in the PCR reactions described below.

25 Specific PCR primers was designed (Table 11) based on the sequences of the rat GABA<sub>B</sub> receptor 1a and 1b cDNA and human GABA<sub>B</sub> receptor 1a and 1b cDNA

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TABLE 11
Primers used for RT-PCR on mRNA from human fetal brain

Nr	Species	Sequence 5'-3'	SEQ ID NO
838	Rat	ATGCGCGCCGGCAGCCAACATGCTGCTGCTGCTGCTGCTCTCTCT	62
863	Rat	GGTCATCCAGCGTTGAGGTGAAGAC	63
864	Rat	GAAGGTTGCCAGATTATACATCCGC	64
865	Rat	CCACGATGATTCGAGCATCTTGACG	65
937	Human	CTGGTTCCTCCCAATGTG	66
1015	Human	CCAGTGGACTATGAGATTGAG	67

cDNA fragments encoding parts of the human GABA<sub>B</sub> receptors were amplified directly by PCR using the designed primers with the generated cDNA molecules as templates. PCR experiments with primers 838, 863,864 and 865 were carried out using the Perkin Elmer Taq DNA polymerase with Gene Amp (Roche Molecular System Inc., NJ, USA) with the following PCR program: +95°C for 1 minute, +50°C for 30 seconds,+72°C for 3 minutes,
 repeated 44 times and finally +72°C for 7 minutes. PCR experiments with primers 937 and 1015 were carried out using the Expand Long Template PCR System (Boehringer Mannheim GmbH, Germany) with the following PCR program: +94°C for 2 minutes; +94°C for 10 seconds, +55°C for 30 seconds, +68°C for 3 minutes repeated 35 times; and finally +68°C for 7 minutes. The primer combinations 838 and 863, 864 and 863, 864 and
 865, 937 and 1015 gave PCR products.

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, WI, USA). The inserts were subjected to nucleotide sequence analysis. The complete nucleotide sequence for all subclones were determined using a ThermoSequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As for primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to the cDNA encoding the GABA<sub>B</sub> receptor were used.

A number of analyzed clones isolated from fetal brain were found to lack 186 base pairs, corresponding to exon 4. This alternative splicing resulted in a cDNA (SEQ ID NO: 70)

encoding a protein (SEQ ID NO: 71) comprising of 899 amino acids and designated GABA<sub>B</sub> receptor 1e.

Other clones from fetal brain lacked 368 base pairs, corresponding to exons 4, 5 and 6, resulting in a cDNA (SEQ ID NO: 72) where a frame shift and a translational stop codon are generated in the sequence corresponding to exon 7, encoding a protein (SEQ ID NO: 73) comprising only 97 amino acids designated GABA<sub>B</sub> receptor 1f.

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One clone lacked 207 base pairs, corresponding to exons 4 and 5, resulting in a cDNA (SEQ ID NO: 74) encoding a protein (SEQ ID NO: 75) comprising 892 amino acids designated GABA<sub>B</sub> receptor 1g.

Another clone had two deletions, the first comprising 186 base pairs corresponding to exon 4, the second comprising 39 base pairs corresponding to part of exon 6, resulting in a cDNA (SEQ ID NO: 76) encoding a protein (SEQ ID NO: 77) comprising 886 amino acids designated GABA<sub>B</sub> receptor 1h.

Yet another clone from adult human brain was found to have a long deletion comprising 1194 base pairs corresponding to base pairs 319 - 1512 of the cDNA encoding human GABA<sub>B</sub> receptor 1a. This deletion corresponds to part of exon 4, exons 5-11, and part of exon 12. This cDNA (SEQ ID NO: 78) encodes a protein (SEQ ID NO: 79) comprising 563 amino acids designated GABA<sub>B</sub> receptor 1i.

One clone isolated from fetal brain lacked 284 base pairs corresponding to part of exon 3

25 and the whole exon 4, generating a frame shift and a translational stop codon in the sequence corresponding to exon 5. This cDNA (SEQ ID NO: 80) encodes a protein (SEQ ID NO: 81) comprising only 105 amino acids designated GABA<sub>B</sub> receptor 1j.

30 EXAMPLE 8. Analysis of cDNA encoding human GABA<sub>B</sub> receptors from Jurkat cells

A guanidinisothiocyanate/CsCl method was used to isolate total RNA from Jurkat cells. First strand cDNA synthesis reactions were performed using the first strand cDNA

Synthesis kit from Amersham Pharmacia Biotech (Uppsala, Sweden). The pd(N)<sub>6</sub> primer was used to prime the first-strand synthesis. The generated cDNA molecules were used as templates in the PCR reactions described below.

5 Specific PCR primers was designed (Table 12) based on the sequences of the human GABA<sub>B</sub> receptor 1a and 1b cDNA

TABLE 12
Primers used for RT-PCR on mRNA from Jurkat cells

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product.

Nr	Species	Sequence 5'-3'	SEQ ID NO
937	Human	CTGGTTCCTCCCAATGTG	68
1015	Human	CCAGTGGACTATGAGATTGAG	69

cDNA fragments encoding parts of the human GABA<sub>B</sub> receptors were amplified directly by PCR using the designed primers with the generated cDNA molecules as templates. The PCR experiment was carried out using the Expand Long Template PCR System (Boehringer Mannheim GmbH, Germany) with the following PCR program: +94°C for 2 minutes; +94°C for 10 seconds, +55°C for 30 seconds, +68°C for 3 minutes repeated 35 times; and finally +68°C for 7 minutes. The primer combination 937 and 1015 gave a PCR

The PCR products were subcloned into the pGEM-T vector from Promega (Madison, WI, USA). The inserts were subjected to nucleotide sequence analysis. The complete nucleotide sequence for all subclones were determined using a ThermoSequenase dye terminator cycle sequencing pre-mix kit (Amersham Pharmacia Biotech, Uppsala, Sweden). As for primers for sequencing reactions specific oligonucleotides complementary to the vector pGEM-T or primers complementary to the cDNA encoding the GABA<sub>B</sub> receptor were used.

Two clones had two deletions, the first comprising 368 base pairs corresponding to exons 4, 5 and 6, the second comprising 151 base pairs corresponding to exon 15 where a frame shift and a translational stop codon are generated in the sequence corresponding to exon 7. This cDNA (SEQ ID NO: 82) encodes a protein (SEQ ID NO: 83) comprising only 98

amino acids designated GABA<sub>B</sub> receptor 1k which is identical to the GABA<sub>B</sub> receptor 1f described above.

Another two clones also had two deletions, the first comprising 246 base pairs corresponding to part of exon 4, exon 5 and exon 6, the second comprising 149 base pairs corresponding to exon 15, generating a frame shift and a translational stop codon in the sequence corresponding to exon 16. This cDNA (SEQ ID NO: 84) encodes a protein (SEQ ID NO: 5) comprising 496 amino acids designated GABA<sub>B</sub> receptor 11.

Additional variants of cDNA encoding the human GABA<sub>B</sub> receptors can be identified in a similar manner using PCR primers based on the sequence of the cDNA and genomic fragments encoding the human GABA<sub>B</sub> receptors disclosed in the present application.

The biological activity of these variants of the human GABA<sub>B</sub> receptor can be evaluated by transfection of suitable host cells with expression vectors comprising the corresponding cDNA sequences and subsequent measurement of binding of labeled ligands or activation of the receptor or modulation of receptor function.

#### 20 EXAMPLE 9: Generation of antibodies

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Antibodies were raised in rabbits against four different BSA-conjugated 20 amino acids long synthetic peptides corresponding to selected regions of the human and canine GABA<sub>B</sub> receptor extracellular domain. Two were directed against a sequence common for GABA<sub>B</sub> receptor 1a and b (ab1 and ab2), one against a GABA<sub>B</sub> receptor 1a-specific region (a1) and one against a GABA<sub>B</sub> receptor 1b-specific sequence (b1). To allow BSA-conjugation, a cystein residue was added to the N-terminus in all peptides but a1, which contains an endogenous cystein.

30 Peptide al: (SEQ ID NO: 49 amino acids 18-37)

Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile Ile His Pro Pro Trp

Peptide ab1: (SEQ ID NO: 49 amino acids 197-216 with N-terminally added Cys)

Cys - Glu Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu Ile

His His

5 Peptide ab2: (SEQ ID NO: 49 amino acids 271-290 with N-terminally added Cys)

Cys - Ser Pro Ala Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro

Ser Ala

Peptide b1: (SEQ ID NO: 57 amino acids 30-47 with N-terminally added Cys)

Cys - Ser His Ser Pro His Leu Pro Arg Pro His Ser Arg Val Pro Pro His Pro Ser

The antibodies were purified from rabbit serum by affinity chromatography using the corresponding immobilized peptide and subsequently used to detect expression of recombinant GABA<sub>B</sub> receptor isoforms on Western blots.

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EXAMPLE 10: Heterologous expression of GABA<sub>B</sub> receptor isoforms in mammalian cells

A HindIII / SalI cDNA fragment encoding the human GABA<sub>B</sub> receptor 1b isoform was cloned into a BPV (bovine papilloma virus)-based expression vector containing the mMT-1 (murine metallothioneine) promoter. Using the calcium phosphate method, murine C127 cells were co-transfected with the GABA<sub>B</sub> receptor expression construct and an expression plasmid containing a G418 resistance marker gene using the calcium phosphate method. G418 resistant clones were evaluated by Western blot analysis for expression of the approximately 100 kDa GABA<sub>B</sub> receptor 1b isoform(Figure 2). Similarly, but using Lipofectamine (Life Technologies Inc, Rockville, MD, USA) for transfection, the human GABA<sub>B</sub> receptor 1b isoform was expressed in human HEK-293 cells using the pCI-neo expression vector. The identity of the heterologously expressed receptor was verified in HEK-293 cells by Western blot analysis and radiologand binding experiments.

30

A cDNA fragment encoding the human GABA<sub>B</sub> receptor 1d isoform was cloned into a BPV-based expression vector containing the mMT-1 promotor. Using the calcium phosphate method, murine C127 cells were co-transfected with the GABA<sub>B</sub> receptor

expression construct and an expression plasmid containing a G418 resistance marker gene. G418 resistant clones, and concentrated medium from the same clones, were evaluated for GABA<sub>B</sub> receptor 1d isoform expression by Western blot analysis (Figure 3). The experiment revealed the human GABA<sub>B</sub> receptor 1d to be a secreted isoform.

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EXAMPLE 11: Heterologous expression of GABA<sub>B</sub> receptor isoforms in E. coli

A cDNA fragment encoding the human GABA<sub>B</sub> receptor 1d isoform was cloned into a modified pET (Pharmacia Amersham, Uppsala, Sweden) vector downstream of a STII (heat stable enterotoxin II of *E. coli*) signal peptide and followed by a thrombin clevage site and a hexahistidine tag. The expression construct was subsequently used to transform the BL21 (DE3) *E. coli* strain. Western blot analysis of IPTG-induced bacteria revealed expression of a human GABA<sub>B</sub> receptor 1d isoform of expected size(Figure 4).

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In addition, the human GABA<sub>B</sub> receptor 1d isoform was successfully expressed without fusion to a bacterial signal peptide in *E. coli* strain AD494 (DE3).

20 EXAMPLE 12: Method for the screening of substances which are GABA<sub>B</sub> receptor antagonists or agonists

GABA<sub>B</sub> receptor expressing cells, transgenic animals or cells and tissues derived thereof, are used to screen substance libraries for antagonist or agonist activities. Screening is performed as ligand binding assays or functional assays. For screening, cells and tissues are prepared in various ways, each uniquely suited to its purpose. Ligand binding assays are performed *in vivo* or *in vitro*. Functional assays exemplified by, but not limited to, Ca<sup>++</sup>-responses, cAMP-responses and effects on Cl<sup>-</sup> and K<sup>+</sup> channels, are performed in living cells, broken cells, isolated cell membranes, as well as in tissues and in living animals.

## **CLAIMS**

 A nucleic acid molecule encoding a human or canine GABAB receptor, or a functionally equivalent modified form thereof.

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- A nucleic acid molecule according to claim 1 encoding a human GABAB receptor, or a functionally equivalent modified form thereof.
- A nucleic acid molecule according to claim 2 encoding the human GABAB receptor
   1a, or a functionally equivalent modified form thereof.
  - 4. A nucleic acid molecule according to claim 3 selected from:
    - (a) a nucleic acid molecule comprising a nucleotide sequence set forth as SEQ ID NO: 48;
- (b) a nucleic acid molecule comprising a nucleotide sequence capable of hybridizing, under stringent hybridization conditions, to a nucleotide sequence complementary to the polypeptide coding region of a DNA molecule as defined in (a); and
  (c) a nucleic acid molecule comprising a nucleotide sequence which is degenerate as a result of the genetic code to a nucleotide sequence as defined in (a) or (b).

- A nucleic acid molecule according to claim 2 encoding the human GABAB receptor
   lb, or a functionally equivalent modified form thereof..
- 6. A nucleic acid molecule according to claim 5 selected from:
- (a) nucleic acid molecules comprising a nucleotide sequence set forth as SEQ ID NO:50;
  - (b) nucleic acid molecules comprising a nucleotide sequence capable of hybridizing, under stringent hybridization conditions, to a nucleotide sequence complementary the polypeptide coding region of a DNA molecule as defined in (a); and
- 30 (c) nucleic acid molecules comprising a nucleotide sequence which is degenerate as a result of the genetic code to a nucleotide sequence as defined in (a) or (b).

- 7. A nucleic acid molecule according to claim 2 encoding the human GABAB receptor

  1c, or a functionally equivalent modified form thereof.
- 8. A nucleic acid molecule according to claim 7 selected from:
- 5 (a) nucleic acid molecules comprising a nucleotide sequence set forth as SEQ ID NO: 54;
  - (b) nucleic acid molecules comprising a nucleotide sequence capable of hybridizing, under stringent hybridization conditions, to a nucleotide sequence complementary the polypeptide coding region of a DNA molecule as defined in (a); and
- 10 (c) nucleic acid molecules comprising a nucleotide sequence which is degenerate as a result of the genetic code to a nucleotide sequence as defined in (a) or (b).
  - A nucleic acid molecule according to claim 2 encoding the human GABAB receptor
     1d, or a functionally equivalent modified form thereof..

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- 10. A nucleic acid molecule according to claim 9 selected from:
  - (a) nucleic acid molecules comprising a nucleotide sequence set forth as SEQ ID NO: 56;
- (b) nucleic acid molecules comprising a nucleotide sequence capable of hybridizing,
  under stringent hybridization conditions, to a nucleotide sequence complementary the
  polypeptide coding region of a DNA molecule as defined in (a); and
  (c) nucleic acid molecules comprising a nucleotide sequence which is degenerate as a
  - result of the genetic code to a nucleotide sequence as defined in (a) or (b).
- 25 11. A nucleic acid molecule according to claim 1 encoding a canine GABAB receptor, or a functionally equivalent modified form thereof.
  - A nucleic acid molecule according to claim 11 encoding the canine GABAB receptor
     1a, or a functionally equivalent modified form thereof.

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13. A nucleic acid molecule according to claim 12 selected from

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- (a) nucleic acid molecules comprising a nucleotide sequence set forth as SEQ ID NO: 52;
- (b) nucleic acid molecules comprising a nucleotide sequence capable of hybridizing, under stringent hybridization conditions, to a nucleotide sequence complementary the polypeptide coding region of a DNA molecule as defined in (a); and
- (c) nucleic acid molecules comprising a nucleotide sequence which is degenerate as a result of the genetic code to a nucleic acid sequence as defined in (a) or (b).
- A nucleic acid molecule according to claim 11 encoding the canine GABAB receptor 10 lc, or a functionally equivalent modified form thereof.
  - 15. A nucleic acid molecule according to claim 14 selected from(a) nucleic acid molecules comprising a nucleotide sequence set forth as SEQ ID NO:58;
- (b) nucleic acid molecules comprising a nucleotide sequence capable of hybridizing, under stringent hybridization conditions, to a nucleotide sequence complementary the polypeptide coding region of a DNA molecule as defined in (a); and (c) nucleic acid molecules comprising a nucleotide sequence which is degenerate as a result of the genetic code to a nucleic acid sequence as defined in (a) or (b).

16. A recombinant polypeptide encoded by a nucleotide sequence according to any one of claims 1 to 15.

- 17. The polypeptide according to claim 16 comprising an amino acid sequence set forth as SEQ ID NO: 49, 51, 53, 55, 57 or 59 in the Sequence Listing.
  - 18. A polypeptide according to claim 16 which has been posttranslationally modified.
- 19. A vector transformed with a nucleic acid molecule according to any one of claims 130 to 15.
  - 20. A cultured host cell harboring a vector according to claim 19.

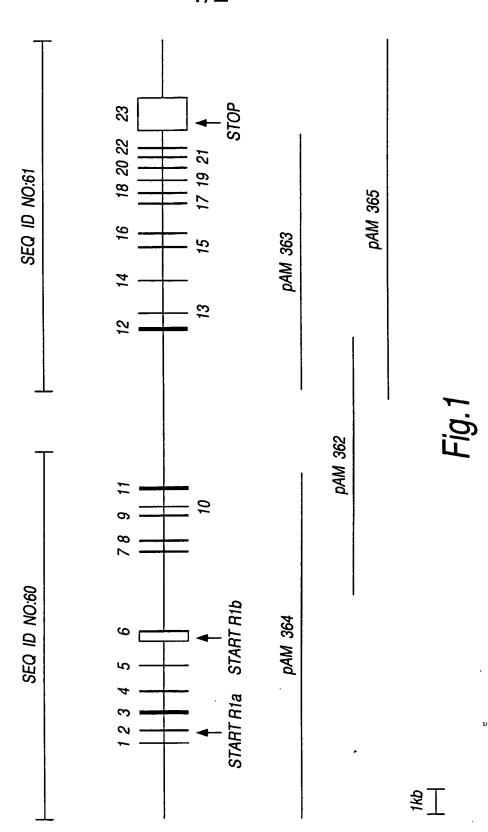
21. A process for the production of a GABAB receptor, said process comprising culturing a host cell according to claim 20 under conditions whereby the said polypeptide is produced, and recovering the said polypeptide.

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- 22. A method for the screening of compounds which are inhibitors of transient lower esophageal sphincter relaxations, said method comprising the use of a nucleic acid molecule encoding a GABAB receptor.
- 10 23. A method according to claim 22 wherein the said nucleic acid molecule encoding a GABAB receptor is the nucleic acid molecule according to any one of claims 1 to 15.
  - 24. A method for the screening of compounds which are agonists or antagonists to a GABAB receptor, said method comprising the use of a nucleic acid molecule according to any one of claims 1 to 15.
  - 25. A method according to any one of claims 22 to 24 comprising the steps (a) transforming a cultured cell with a nucleic acid molecule encoding a GABAB receptor, so that a GABAB receptor is expressed on the surface of the cell; (b) contacting a test compound with the said cell; and (c) determining whether the test compound binds to, and/or activate, the GABAB receptor.
  - 26. A pharmaceutical compositions comprising a soluble GABA<sub>B</sub> receptor further comprising one or more pharameeutical acceptable carriers and/or diluents.

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SUBSTITUTE SHEET (RULE 26)

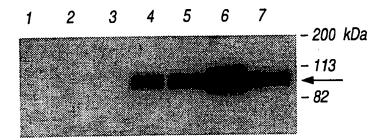


Fig.2

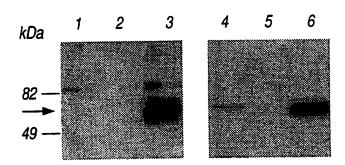


Fig.3

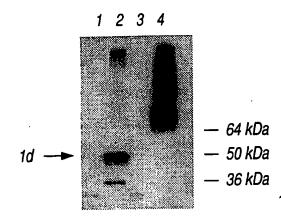


Fig.4

SUBSTITUTE SHEET (RULE 26)

SEQUENCE LISTING

5

15

- (1) GENERAL INFORMATION:
  - (i) APPLICANT:
    - (A) NAME: ASTRA AB
- 10 (B) STREET: Västra Mälarehamnen 9
  - (C) CITY: Södertälje
  - (E) COUNTRY: Sweden
  - (F) POSTAL CODE (ZIP): S-151 85
  - (G) TELEPHONE: +46-8-553 260 00
  - (H) TELEFAX: +46-8-553 288 20
  - (I) TELEX: 19237 astra s
  - (ii) TITLE OF INVENTION: New nucleotide sequences
- 20 (iii) NUMBER OF SEQUENCES: 85
  - (iv) COMPUTER READABLE FORM:
    - (A) MEDIUM TYPE: Floppy disk
    - (B) COMPUTER: IBM PC compatible
  - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
    - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30 (EPO)
  - (2) INFORMATION FOR SEQ ID NO: 1:

30

25

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 27 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
- 35 (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

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GTTTCTTCTC GGATCCAGCT GTGCCTG

27

H1865-1 WO SEQ

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5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 28 base pairs</li><li>(B) TYPE: nucleic acid</li><li>(C) STRANDEDNESS: single</li><li>(D) TOPOLOGY: linear</li></ul>	
	(ii) MOLECULE TYPE: DNA	
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40	(ii) MOLECULE TYPE: DNA	
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	ATGCGCGCCG GCAGCCAACA TGCTGCTGCT GCTGCTGGTG C	. 41
5	(2) INFORMATION FOR SEQ ID NO: 5:	
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	(A) LENGTH: 46 base pairs	
	(B) TYPE: nucleic acid	
10	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:	
	CGGTCGACTC ACTTGTAAAG CAAATGTACT CGACTCCCAT CACAGC	46
. 20	(2) INFORMATION FOR SEQ ID NO: 6:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 50 base pairs	
	(B) TYPE: nucleic acid	
25	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:	•
	ATGCGCGCCG GCAGCCAACA TGCTGCTGCT GCTGCTGGTG CCTCTCTTCC	50
35	(2) INFORMATION FOR SEQ ID NO: 7:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 40 base pairs	

(ii) MOLECULE TYPE: DNA

(B) TYPE: nucleic acid(C) STRANDEDNESS: single(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7: CAGGCACAGC TGGATCCGAG AAGAAACTCT GTCGGAAAGT 40 5 (2) INFORMATION FOR SEQ ID NO: 8: (i) SEQUENCE CHARACTERISTICS: 10 (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8: GGTCATCCAG CGTTGAGGTG AAGAC 25 20 (2) INFORMATION FOR SEQ ID NO: 9: (i) SEQUENCE CHARACTERISTICS: 25 (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9: GAAGGTTGCC AGATTATACA TCCGC 25 35 (2) INFORMATION FOR SEQ ID NO: 10: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear H1865-1 WO SEQ

	(ii) MOLECULE TYPE: DNA	
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	(C) STRANDEDNESS: single	
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	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:	
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	(A) LENGTH: 21 base pairs	
	(B) TYPE: nucleic acid	
30	(C) STRANDEDNESS: single (D) TOPOLOGY: linear	•
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	(B) TYPE: nucleic acid	
	H1865-1 WO SEQ 5	

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5	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 27 base pairs</li> <li>(B) TYPE: nucleic acid</li> <li>(C) STRANDEDNESS: single</li> <li>(D) TOPOLOGY: linear</li> </ul>	
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20	(D) TOPOLOGY: linear	
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35	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:	
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	(B) TYPE: nucleic acid		
	(C) STRANDEDNESS: single		
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	(ii) MOLECULE TYPE: DNA		
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	(B) TYPE: nucleic acid		
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	(D) TOPOLOGY: linear		
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               (B) TYPE: nucleic acid
               (C) STRANDEDNESS: single
               (D) TOPOLOGY: linear
25
        (ii) MOLECULE TYPE: DNA
        (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:
    CGTCAAGATG CTCGAATCAT CG
                                                                             22
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35
               (A) LENGTH: 20 base pairs
               (B) TYPE: nucleic acid
               (C) STRANDEDNESS: single
               (D) TOPOLOGY: linear
        (ii) MOLECULE TYPE: DNA
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:
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H1865-1 WO SEQ

CAGGGGGCTC AGAGGGTCCC

10

25

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- 5 (2) INFORMATION FOR SEQ ID NO: 25:
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    - (B) TYPE: nucleic acid
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    - (ii) MOLECULE TYPE: DNA
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CAGGCACAGC TGGATCCGAG AAGAAACTCT GTCGGAAAGT

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- 20 (2) INFORMATION FOR SEQ ID NO: 26:
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    - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
    - (ii) MOLECULE TYPE: DNA
- 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:

CGGTCGACTC ACTTGTAAAG CAAATGTACT CGACTCCCAT CACAGCTAAG

- 35 (2) INFORMATION FOR SEQ ID NO: 27:
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    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA

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20 -	CCACGATGAT TCGAGCATCT TGACG	25
	(2) INFORMATION FOR SEQ ID NO: 29:	
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 24 base pairs</li><li>(B) TYPE: nucleic acid</li><li>(C) STRANDEDNESS: single</li><li>(D) TOPOLOGY: linear</li></ul>	
30	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:	
35	CTACCGCGCA ATGAACTCCT CGTC	24
	(2) INFORMATION FOR SEQ ID NO: 30:	
40	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 24 base pairs</li><li>(B) TYPE: nucleic acid</li><li>(C) STRANDEDNESS: single</li></ul>	,
	(D) TOPOLOGY: linear	

	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:	
5	CGAGGTGGCG TTGGGGGTCT GTGC	24
	(2) INFORMATION FOR SEQ ID NO: 31:	
10	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 27 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: single	
15	(D) TOPOLOGY: linear	
•	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:	
20	CCATCCTAAT ACGACTCACT ATAGGGC	27
	(2) INFORMATION FOR SEQ ID NO: 32:	
25	(a) and a contract of the cont	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 23 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: single	
30	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:	
35	ACTCACTATA GGGCTCGAGC GGC	23
	(2) INFORMATION FOR SEQ ID NO: 33:	
40		
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 20 base pairs  (B) TYPE: nucleic acid	
	H1865-1 WO SEO 12	
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(C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33: GACGCTTATC GAGCAGCTTC 20 10 (2) INFORMATION FOR SEQ ID NO: 34: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA 20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34: AGCCCAGAAC TCACAGGGGG ACAT 24 25 (2) INFORMATION FOR SEQ ID NO: 35: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA 35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35: GCTTCAAGCC AGGTACGAAC TAA 23 · 40 (2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS: H1865-1 WO SEQ

(A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 5 (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGAGCACCCC CAAGCCCCAC TG 22 (2) INFORMATION FOR SEQ ID NO: 37: (i) SEQUENCE CHARACTERISTICS: 15 (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 20 (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37: CTGGTTCCTC CCAATGTG . 18 (2) INFORMATION FOR SEQ ID NO: 38: (i) SEQUENCE CHARACTERISTICS: 30 (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 35 (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38: CCTCTCACTC CCCTCATCTC 20, (2) INFORMATION FOR SEQ ID NO: 39: H1865-1 WO SEQ 14

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	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 22 base pairs	
	(B) TYPE: nucleic acid	
5	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:	
	AAGCCAACCT TCCCTGCTTC TC	22
15	(2) INFORMATION FOR SEQ ID NO: 40:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 18 base pairs	
	(B) TYPE: nucleic acid	
20	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:	
	CTGGTTCCTC CCAATGTG	18
30	(2) INFORMATION FOR SEQ ID NO: 41:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 20 base pairs	
	(B) TYPE: nucleic acid	
35	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:	,
	GACGCTTATC GAGCAGCTTC	20
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	(2) INFORMATION FOR SEQ ID NO: 42:	
5	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 24 base pairs</li> <li>(B) TYPE: nucleic acid</li> <li>(C) STRANDEDNESS: single</li> <li>(D) TOPOLOGY: linear</li> </ul>	
10	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:	
15	CTACCGCGCA ATGAACTCCT CGTC	24
	(2) INFORMATION FOR SEQ ID NO: 43:	
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 24 base pairs</li><li>(B) TYPE: nucleic acid</li><li>(C) STRANDEDNESS: single</li><li>(D) TOPOLOGY: linear</li></ul>	
25	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:	
30	CCTTCTTCTC CTCCTTCTTA GTGA	24
	(2) INFORMATION FOR SEQ ID NO: 44:	
35	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 2883 base pairs</li> <li>(B) TYPE: nucleic acid</li> <li>(C) STRANDEDNESS: single</li> <li>(D) TOPOLOGY: linear</li> </ul>	
40	(ii) MOLECULE TYPE: cDNA to mRNA	1
	(iii) HYPOTHETICAL: NO	

(iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Rattus norwegicus 5 (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1..2883 (x) PUBLICATION INFORMATION: 10 (A) AUTHORS: Kaupmann, K Huggel, K Heid, J Flor, P Bischoff, S 15 Mickel, S McMaster, G Angst, C Bittiger, H Froestl, W 20 (B) TITLE: Expression cloning of GABA-B receptors uncovers similarity to metabotropic glutamate receptors (C) JOURNAL: Nature (D) VOLUME: 386 25 (F) PAGES: 239-246 (G) DATE: 20 march-1997 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44: 48 Met Leu Leu Leu Leu Val Pro Leu Phe Leu Arg Pro Leu Gly Ala 1 5 10 35 GGC GGG GCG CAG ACC CCC AAC GCC ACC TCG GAA GGT TGC CAG ATT ATA 96 Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile Ile 20 25 CAT CCG CCC TGG GAA GGT GGC ATC AGG TAC CGT GGC TTG ACT CGC GAC 144 His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg Asp 35 CAG GTG AAG GCC ATC AAC TTC CTG CCT GTG GAC TAT GAG ATC GAA TAT 192 H1865-1 WO SEQ 17

	Gln	Val	Lys	Ala	Ile	Asn	Phe	Leu	Pro	Val	Asp	Tyr	Glu	Ilė	Glu	Tyr		
		50					55					60						
	GTG	TGC	CGA	GGG	GAG	CGC	GAG	GTG	GTG	GGG	CCC	AAG	GTG	CGC	AAA	TGC		240
5	Val	Cys	Arg	Gly	Glu	Arg	Glu	Val	Val	Gly	Pro	Lys	Val	Arg	Lys	Cys		
	65					70					75					80		
	CTG	GCC	AAC	GGC	TCC	TGG	ACG	GAT	ATG	GAC	ACA	CCC	AGC	CGC	TGT	GTC		288
	Leu	Ala	Asn	Gly	Ser	Trp	Thr	Asp	Met	Asp	Thr	Pro	Ser	Arg	Cys	Val		
10					85					90					95			
	CGA	ATC	TGC	TCC	AAG	TCT	TAT	TTG	ACC	CTG	GAA	AAT	GGG	AAG	GTT	TTC		336
	Arg	Ile	Cys	Ser	Lys	Ser	Tyr	Leu	Thr	Leu	Glu	Asn	Gly	Lys	Val	Phe		
				100					105					110				
15																		
			GGT														•	384
	Leu	Thr	Gly	Gly	Asp	Leu	Pro		Leu	Asp	Gly	Ala	-	Val	Glu	Phe		
			115					120					125					
20	CGA	TGT	GAC	CCC	GAC	TTC	CAT	CTG	GTG	GGC	AGC	TCC	CGG	AGC	GTC	TGT		432
	Arg	Cys	Asp	Pro	Asp	Phe	His	Leu	Val	Gly	Ser	Ser	Arg	Ser	Val	Cys		
		130					135					140						
							100			~~~	010	<b></b>				223		
			GGC															480
25	Ser 145	GIN	Gly	GIN	тър	150	IIII	PIO	гуу	PIO	155	cys	GIN	vaı	ASII	160		
	142					130					133					100		
	ACG	CCA	CAC	TCA	GAA	CGG	CGT	GCA	GTA	TAC	ATC	GGG	GCG	CTG	TTT	CCC		528
	Thr	Pro	His	Ser	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	Ala	Leu	Phe	Pro		
30					165					170					175		•	
	ATG	AGC	GGG	GGC	TGG	CCG	GGG	GGC	CAG	GCC	TGC	CAG	ccc	GCG	GTG	GAG		576
	Met	Ser	Gly	Gly	Trp	Pro	Gly	Gly	Gln	Ala	Cys	Gln	Pro	Ala	Val	Glu		
				180					185					190				
35																		
	ATG	GCG	CTG	GAG	GAC	GTT	AAC	AGC	CGC	AGA	GAC	ATC	CTG	CCG	GAC	TAC		624
	Met	Ala	Leu	Glu	Asp	Val	Asn		Arg	Arg	Asp	Ile	Leu	Pro	Asp	Tyr		
			195					200					205					
40	GAG	СТС	AAG	CTT	ATC	CAC	CAC	GAC	AGC	AAG	TGT	GAC	CCA	GGG	CAA	GCC		672 <sub>v</sub>
	Glu	Leu	Lys	Leu	Ile	His	His	Asp	Ser	Lys	Cys	Asp	Pro	Gly	Gln	Ala		
		210					215		,			220			•			

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	385					390					395					400			
	-	Lys	Tyr	Val	Trp		Leu	Ile	Gly	Trp		Ala	Asp	Asn	Trp	Phe	٠.		
,						TTC									_				00
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	C11	ccc	000	<b>777</b>	Cum	mmm	uh∕-un	CNC	Carc	መልመ	מ מ	CNA	אככ	CTC	արտո	ccc		11	52
35			355					360					365						
	Leu	Lys	Arg	Gln	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	Phe	Tyr	Glu	Thr			
	CTG	AAG	CGT	CAA	GAT	GCT	CGA	ATC	ATC	GTG	GGA	CTT	TTC	TAT	GAG	ACG		11	04
				240					747					550					
30	Thr	Phe	Arg	Gln 340	Ser	Phe	Phe	Ser	Asp 345	Pro	Ala	val	Pro	Val 350	rys	Asn			
						TTC	_				_			_			•	10	56
			-	-	325			_		330			-		335				
دند						Glu													- <del>-</del>
25	ACG	ርጥና	GAT	GAC	CTG	GAG	GAG	CGA	GTG	AAA	GAG	GCT	GGG	ATC	GAG	ATC		10	08
	305					310					315					320			
	Trp	Lys	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr		Glu	Val	Phe	Thr				
	TGG	AAG	AAG	ATC	GCT	ACC	ATC	CAA	CAG	ACC	ACC	GAG	GTC	TTC	ACC	TCA		9	60
20																			
	1110	290	204			-10	295	9		_, 5		300		_, 5	1	1			
						CCC Pro												9	12
	200		ama	<b>C</b>	330	000	3.00	000	CEC	222	CMC	mmc.	<i>C</i> 2 2 2	220	mac.	000			10
15			275					280					285						
	Leu	Ser	Asn	Arg	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	His	Pro	Ser			
	TTG	TCA	AAC	CGA	CAG	CGG	TTT	ccc	ACG	TTC	TTC	CGG	ACG	CAT	CCA	TCC		8	64
				200					203					210					
10	Arg	Met	Trp	Asn 260	Leu	Ile	val	ьeu	Ser 265	ıyr	GIĀ	ser	ser	Ser 270	rro	ATA			
						ATT	_				_					_		8	16
									•										
					245					250					255				
-						Ser													
5	СТС	ATG	ССТ	GGC	TGT	AGT	TCT	GTC	TCC	ACA	CTT	GTA	GCT	GAG	GCT	GCC		7	68
	225					230					235					240			
	Thr	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	_	Pro	Ile	Lys	Ile				
	ACC	AAG	TAC	TTG	TAC	GAA	CTA	CTC	TAC	AAT	GAC	ccc	ATC	AAG	ATĆ	ATT	٠.	7	<del>2</del> 0
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			AAA Lys											•	•		1728
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40	545					550					555					560	
	Ser	Tyr	Lys	Lys	Ile	Gly	Tyr	Tyr	Asp	Ser	Thr	Lys	Asp	Asp	Leu	Ser	
	AGC	TAC	AAG	AAG	ATC	GGC	TAC	TAC	GAC	AGC	ACC	AAG	GAT	GAT	CTT	TCC	1680
,,	Ser	530	561	**** 9	11C C		535		204		<b>014</b>	540	254	· · ·	,	J-1	
35																Gly	
	<b>У</b> СС	GGC	TCC	ርርር የ	ልጥር	GC 2	TYCG	ልሮኔ	СФФ	ልሞሮ	GAG	CAG	ርሞን	CAG	GGC	GGC	1632
			515					520					525			•	
	Asn	Ser	Ser	Ser	Phe	Glu	Gly	Val	Ser	Gly	His	Val	Val	Phe	Asp	Ala	
30	AAC	TCC	TCC	TCC	TTT	GAG	GGC	GTT	TCT	GGC	CAT	GTG	GTC	TTT	GAT	GCC	1584
	* 11C	1,511	-1-	500			~ * * *		505				- 7 -	510		<b>-</b>	
			Tyr														1000
25	יניתותי	λÁΩ	TAC	ልልሮ	ממ	CAG	ACC	<u>አ</u> ጥጥ	ארא	GYC	CAG	ልጥሮ	ጥልሮ	ന്ദര	GCC	ATG	1536
25					485					490					495		
	Asn	Lys	Thr	Ser	_	Gly	Gly	Gly	Arg		Gly	Val	Arg	Leu		Asp	
			ACG														1488
20	465					470	-			-	475					480	
			Pro														
	GAG	GCA	CCA	CTG	GCC	TAT	GAT	GCT	ATC	TGG	GCC	TTG	GCT	TTG	GCC	TTG	1440
		450					455				•	460				•	
15	Leu		Lys	Arg	Leu	Lys		His	Pro	Glu	Glu		Gly	Gly	Phe	Gln	
			AAG														1392
			435					440					445			_	
••			Arg														
10	AAC	ACC	CGA	AGC	ATT	TCC	AAC	ATG	ACG	TCA	CAG	GAA	ттт	GTG	GAG	AAA	1344
				420					425					430			
	Ala	Val	Glu		His	Ile	Thr	Thr		Ile	Val	Met	Leu		Pro	Ala	
			GAG														1296
5																	
					405					410					415		
			Tyr													•	
	AAG	ACC	TAT	GAC	CCG	TCA	ATC	ААТ	TGT	ACA	GTG	GAA	GÄA	ATG	ACC	GAĞ	1248
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					565					570		•	•	٠	575		••	
	ACC	ጥጥር	ርጥር	ልጥሮ	AAG	ACA	ጥጥር	CGT	ттс	CTG	ጥርጥ	CAG	ΔΑΔ	רתי	TTT	ልጥሮ		1776
															Phe	_		17.0
5	****	Deu	V 44 1	580	Lys				585	204	502	· · · ·	2,5	590	1110	776		
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	TCC	GTC	тса	GTT	СТС	TCC	AGC	CTG	GGC	АТТ	GTT	СТТ	GCT	GTT	GTC	тст		1824
•															Val			
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10																		
	CTG	TCC	TTT	AAC	ATC	TAC	AAC	TCC	CAC	GTT	CGT	TAT	ATC	CAG	AAC	TCC		1872
										_					Asn			
		610				_	615				_	620						
15	CAG	ccc	AAC	CTG	AAC	AAT	CTG	ACT	GCT	GTG	GGC	TGC	TCA	CTG	GCA	CTG		1920
	Gln	Pro	Asn	Leu	Asn	Asn	Leu	Thr	Ala	Val	Gly	Cys	Ser	Leu	Ala	Leu		
	625					630					635					640		
	GCT	GCT	GTC	TTC	CCT	CTC	GGG	CTG	GAT	GGT	TAC	CAC	ATA	GGG	AGA	AGC		1968
20	Ala	Ala	Val	Phe	Pro	Leu	Gly	Leu	Asp	Gly	Tyr	His	Ile	Gly	Arg	Ser		
	•				645					650					655			
	CAG	TTC	CCG	TTT	GTC	TGC	CAG	GCC	CGC	CTT	TGG	CTC	TTG	GGC	TTG	GGC		2016
	Gln	Phe	Pro	Phe	Val	Cys	Gln	Ala	Arg	Leu	Trp	Leu	Leu	Gly	Leu	Gly		
25				660					665					670				
	TTT	AGT	CTG	GGC	TAT	GGC	TCT	ATG	TTC	ACC	AAG	ATC	TGG	TGG	GTC	CAC		2064
	Phe	Ser	Leu	Gly	Tyr	Gly	Ser	Met	Phe	Thr	Lys	Ile	Trp	Trp	Val	His		
			675					680					685					
30																		
					AAG											CTA		2112
	Thr		Phe	Thr	Lys	Lys		Glu	Lys	Lys	Glu	_	Arg	Lys	Thr	Leu		
		690					695					700						
						m												
35															ATG			2160
		Pro	Trp	Lys	Leu	_	Ala	Thr	Val	GIY		Leu	Val	Gly	Met	_		
	705					710					715					720		
		ama	<b>.</b>	omm.	222	<b>&gt;</b> ma	maa	~~~	3 MM	omo	a. a		mma	a.a	001			2200
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40	Val	Leu	Thr	Leu		ire	ттр	GIN	TIE		Asp	Pro	Leu	HIS	Arg	Thr		ν
					725					730					735			
		<b>636</b>	a com	Mun	000	N N C	C2.C	CAA	003	220	CA A	C2.C	አመጣ	Cym	CMC	maa		2256
	ATT'	GAG	ACT	.T.T.T.	GCC	AAG	GAG	GAA	CCA	AAG	GAA	GAC	ATC	GAT	GTC	TCC	•	2256
	77.5	<i></i>	1110	OFC														
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	Ile	Glu	Thr		Ala	Lys	Glu	Glu		Lys	Glu	Asp	Ile		Val	Ser	
				740					745					750			
	ATT	CTG	CCC	CAG	TTG	GAG	CAC	TGC	AGC	TCC	AAG	AAG	ATG	AAT	ACG	TGG	2304
5	Ile	Leu	Pro	Gln	Leu	Glu	His	Суз	Ser	Ser	Lys	Lys	Met	Asn	Thr	Trp	
			755					760					765				
						_						CTG					2352
	Leu	_	Ile	Phe	Tyr	Gly	_	Lys	Gly	Leu	Leu	Leu	Leu	Leu	GIY	Ile	
10		770					775					780					
	ттт	CTT	GCT	TAC	GAA	ACC	AAG	AGC	GTG	TCC	ACT	GAA	AAG	ATC	ААТ	GAC	2400
	Phe	Leu	Ala	Tyr	Glu	Thr	Lys	Ser	Val	Ser	Thr	Glu	Lys	Ile	Asn	Asp	
	785					790					795					800	
15																	
	CAC	AGG	GCC	GTG	GGC	ATG	GCT	ATC	TAC	AAT	GTC	GCG	GTC	CTG	TGT	CTC	2448
	His	Arg	'Ala	Val	_	Met	Ala	Ile	Tyr		Val	Ala	Val	Leu	-	Leu	
					805					810					815		
20	АТС	АСТ	GCT	ССТ	GTG	ACC	ATG	ATC	CTT	TCC	AGT	CAG	CAG	GAC	GCA	GCC	2496
												Gln					
				820					825					830			
	TTT	GCC	TTT	GCC	TCT	CTG	GCC	ATC	GTG	TTC	TCT	TCC	TAC	ATC	ACT	CTG	2544
25	Phe	Ala		Ala	Ser	Leu	Ala		Val	Phe	Ser	Ser	_	Ile	Thr	Leu	
			835					840					845				
	CUV	CTC	ርሞር	անարան	GTG.	רככ	AAG	ΔΤΩ	CGC	AGG	ርጥር	ATC	ACC	CGA	GGG	GAA	2592
												Ile					
30		850					855					860		-	-		
																·	
	TGG	CAG	TCT	GAA	ACG	CAG	GAC	ACC	ATG	AAA	ACA	GGA	TCA	TCC	ACC	AAC	2640
	Trp	Gln	Ser	Glu	Thr	Gln	Asp	Thr	Met	Lys	Thr	Gly	Ser	Ser	Thr	Asn	
	865					870					875					880	
35	220		C) C	C	C 2 C	220	maa	CC 3	CMC	mma	CNC	220	<i>-</i>	220	CCA	CAA	2688
												AAG Lys					2000
	ASII	VOII	GIU	Giu	885	nys	Sei	Arg	Dea	890	Giu	пуэ	Giu	VOII	895	GIU	
40	CTG	GAA	AAG	ATC	ATC	GCT	GAG	AAA	GAG	GAG	CGC	GTC	TCT	GAA	CTG	CGC	2736 <sub>v</sub>
	Leu	Glu	Lys	Ile	Ile	Ala	Glu	Lys	Glu	Glu	Arg	Val	Ser	Glu	Leu	Arg	
				900					905					910	•		

CAT CAG CTC CAG TCT CGG CAG CAA CTC CGC TCA CGG CGC CAC CCC CCA 2784 His Gln Leu Gln Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro Pro ACA CCC CCA GAT CCC TCT GGG GGC CTT CCC AGG GGA CCC TCT GAG CCC 2832 Thr Pro Pro Asp Pro Ser Gly Gly Leu Pro Arg Gly Pro Ser Glu Pro 930 935 940 CCT GAC CGG CTT AGC TGT GAT GGG AGT CGA GTA CAT TTG CTT TAC AAG 2880 10 Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr Lys 960 950 955 2883 TGA 15 (2) INFORMATION FOR SEQ ID NO: 45: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 960 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 45: Met Leu Leu Leu Leu Val Pro Leu Phe Leu Arg Pro Leu Gly Ala , 15 1 Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile Ile 20 His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg Asp 40 35 Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu Tyr 50 Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys Cys 70 Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys Val

85 90 95

Arg Ile Cys Ser Lys Ser Tyr Leu Thr Leu Glu Asn Gly Lys Val Phe 100 105 110

Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala Arg Val Glu Phe
115 120 125

Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser Arg Ser Val Cys
10 130 135 140

Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe Pro 165 170 175

Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln Pro Ala Val Glu 180 185 190

Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr
195 200 205

Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala 25 210 215 220

Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile 225 230 235 240

Leu Met Pro Gly Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala 245 . 250 . 255

Arg Met Trp Asn Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala 260 265 270

Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser 275 280 285

Ala Thr Leu His Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly 290 295 300

Trp Lys Lys Ile Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr Ser 305 310 315 320

20

Thr Leu Asp Asp Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile Thr Phe Arg Gln Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Thr Tyr Asp Pro Ser Ile Asn Cys Thr Val Glu Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly 

Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ilé Gly Arg Ser Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly 25 Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Lys Lys Met Asn Thr Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Gly Ile

770 775 780

Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp 785 790 795 800

His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu 805 810 815

Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala 10 820 825 830

Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu 835 840 845

Val Val Leu Phe Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu 850 855 860

Trp Gln Ser Glu Thr Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn 865 870 875 880

Asn Asn Glu Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg Glu
885 890 895

Leu Glu Lys Ile Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu Arg
25 900 905 910

His Gln Leu Gln Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro Pro 915 920 925

Thr Pro Pro Asp Pro Ser Gly Gly Leu Pro Arg Gly Pro Ser Glu Pro 930 935 940

Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr Lys 945 950 955 960

(2) INFORMATION FOR SEQ ID NO: 46:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2538 base pairs

(B) TYPE: nucleic acid(C) STRANDEDNESS: single(D) TOPOLOGY: linear

35

```
(ii) MOLECULE TYPE: cDNA to mRNA
       (iii) HYPOTHETICAL: NO
        (iv) ANTI-SENSE: NO
        (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Rattus norwegicus
10
        (ix) FEATURE:
              (A) NAME/KEY: CDS
               (B) LOCATION:1..2532
         (x) PUBLICATION INFORMATION:
15
               (A) AUTHORS: Kaupmann, K
                            Huggel, K
                            Heid, J
                            Flor, P
                            Bischoff, M
20
                            Mickel, S
                            McMaster, G
                            Angst, C
                            Bittiger, H
                            Froestl, W
25
               (B) TITLE: Expression cloning of GABA-B receptors
                     uncovers similarity to metabotropic glutamate
                     receptors
               (C) JOURNAL: Nature
               (D) VOLUME: 386
30
               (F) PAGES: 239-246
               (G) DATE: 20 March-1997
        (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:
35
                                                                             48
    ATG GGC CCG GGG GGA CCC TGT ACC CCA GTG GGG TGG CCG CTG CCT CTT
    Met Gly Pro Gly Gly Pro Cys Thr Pro Val Gly Trp Pro Leu Pro Leu
                       5
                                                               15
      1
                                          10
                                                                             96.,
    CTG CTG GTG ATG GCG GCT GGG GTG GCT CCG GTG TGG GCC TCT CAC TCC
    Leu Leu Val Met Ala Ala Gly Val Ala Pro Val Trp Ala Ser His Ser
                                                           30
                 20
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		a	ome.	000	acc	CCM	CAC	ccc	200	cmc	ccc	ccic	cic	ccc	mcc.	መድ አ		144
										_	Pro							Taa
	PIO	HIS	35	PIO	ALG	FIU	nis	40	ALG	Vai	FIU	FIU	45	FIU	Ser	Jer	•	
			,,					40					4.5					
5	GAA	CGG	CGT	GCA	GTA	TAC	ATC	GGG	GCG	CTG	TTT	ccc	ATG	AGC	GGG	GGC		192
-											Phe							
		50				-	55	-				60			-	-		
	TGG	CCG	GGG	GGC	CAG	GCC	TGC	CAG	ccc	GCG	GTG	GAG	ATG	GCG	CTG	GAG		240
10	Trp	Pro	Gly	Gly	Gln	Ala	Cys	Gln	Pro	Ala	Val	Glu	Met	Ala	Leu	Glu		
	65					70					75					80		
	GAC	GTT	AAC	AGC	CGC	AGA	GAC	ATC	CTG	CCG	GAC	TAC	GAG	CTC	AAG	CTT		288
	Asp	Val	Asn	Ser	Arg	Arg	Asp	Ile	Leu	Pro	Asp	Tyr	Glu	Leu	Lys	Leu		
15					85					90					95			
	ATC	CAC	CAC	GAC	AGC	AAG	TGT	GAC	CCA	GGG	CAA	GCC	ACC	AAG	TAC	TTG		336
	Ile	His	His	Asp	Ser	Lys	Cys	Asp	Pro	Gly	Gln	Ala	Thr	Lys	Tyr	Leu		
				100					105					110				
20																		
											ATC							384
	Tyr	Glu		Leu	Tyr	Asn	Asp		Ile	Lys	Ile	Ile		Met	Pro	Gly		
			115					120					125					
					maa		c.mm	C	COM	<b>63.6</b>	00m	000	000	3.mc	mcc	220		432
25											GCT Ala							432
	Cys	130	ser	vaı	Ser	1111	135	Val	ATG	Gru	AIA	140	ALG	Mec	TTD	Non		
		130					133					140						
	Cuhu	Δጣጥ	GTG	ርጥር	TCA	тат	GGC	TCC	AGT	TCA	CCA	GCC	TTG	TCA	AAC	CGA	-	480
30											Pro							
50	145					150					155					160		
	CAG	CGG	TTT	CCC	ACG	TTC	TTC	CGG	ACG	CAT	CCA	TCC	GCC	ACA	CTC	CAC		528
	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	His	Pro	Ser	Ala	Thr	Leu	His		
35					165					170					175			
	AAT	ccc	ACC	CGG	GTG	AAA	CTC	TTC	GAA	AAG	TGG	GGC	TGG	AAG	AAG	ATC		576
	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe	Glu	Lys	Trp	Gly	Trp	Lys	Lys	Ile		
				180					185					190				
40																		υ
											ACC							624
	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	Val	Phe	Thr	Ser	Thr	Leu	Asp	Asp		
			195					200					205				•	
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	CTG	GAG	GAG	CGA	GTG	AAA	GAG	GCT	GGG	ATC	GAG	ATC	АСТ	TTC	CGA	CAG		672
	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala	Gly	Ile	Glu	Ile	Thr	Phe	Ara	Gln		
		210				-	215		-			220						
5																		
•	AGT	שידירי	TTC	TCG	GAT	CCA	GCT	GTG	ССТ	СТТ	AAA	AAC	СТС	AAG	ССТ	CAA		720
			Phe															
	225					230					235			-,, 0		240		
	223					250					200							
10	СУТ	ርርጥ	CGA	ልጥሮ	ልጥር	GTG	GGA	Cuhu	ጥጥር	ጥልጥ	GAG	ACG	GAA	GCC	CGG	222		768
.0			Arg			_	_				_			_				
	лэр	A10	ALG	110	245	Vul	013	Deu		250	014	****			255	шуз		
					243					250					233			
•	COO	מימימי	TGT	CAC	CTC	יימיי	AAG	GAA	AGG	כיייכ	արդուր	ccc	AAG	AAG	ሞልር	CTC		816
15			Cys															010
15	vai	FILE	Cys	260	Vai	171	Lys	GIU	265	Deu	THE	Gry	цуз	270	TYT	Vai		
				200					203					270				
	mcc.	ጥጥር	CTC	ልጥሮ	GGG	TCC	ጥልጥ	GCT	GAC	220	TGG	ጥጥር	AAG	ACC	ጥልጥ	GAC		864
			Leu															001
20	IIP	riie	275	116	GIJ	115	-3-	280	nsp	11311			285		-1-	пор		
20			2,3					200					200					
	୯୯୯	тса	ATC	ААТ	ጥርጥ	ACA	GTG	GAA	GAA	ATG	ACC	GAG	GCG	GTG	GAG	GGC		912
			Ile															
	110	290			4,5		295	0				300				<b>U</b> -3		
25																		
	CAC	ATC	ACC	ACG	GAG	АТТ	GTC	ATG	CTG	AAC	ССТ	GCC	AAC	ACC	CGA	AGC		960
			Thr				_					_		_				
	305					310					315				3	320		
	505																	
30	ልጥጥ	TCC	AAC	ATG	ACG	TCA	CAG	GAA	ттт	GTG	GAG	AAA	СТА	ACC	AAG	CGG		1008
50			Asn				_	_		_	_							
		001			325					330		-,-			335	5		
					323													
	CAC	ΔΑΔ	AGA	CAC	CCC	GAG	GAG	ACT	GGA	GGC	TTC	CAG	GAG	GCA	CCA	CTG		1056
35			Arg															
<i>J J</i>	200	2,0	9	340					345	3				350				
				3.10														
	GCC	ጥልጥ	GAT	GCT	ልጥሮ	TGG	GCC	TTG	GCT	TTG	GCC	<b>ፐፐ</b> G	AAC	AAG	ACG	тст		1104
			Asp															
40	AIG	TYL	355	AIG	116	110	niu	360	niu	200		Lou	365	2,5		501		
40			د د د					200					503					Ľ
	CCA	GC A	GGT	CCm	CCm	ጥርር	CCC	CTC	רפר	ריזים	GAG	GAC	արդուր	AAC	TAC	AAC		1152
																Asn		
	СТА	GTĀ	GTĀ	GTĀ	wrd	Ser	GIY	val	vrā	Jeu	GIU	rap	1116	VOII	- Y -	******	•	

AAC CAG ACC ATT ACA GAC CAG ATC TAC CGG GCC ATG AAC TCC TCC TCC Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser TTT GAG GGC GTT TCT GGC CAT GTG GTC TTT GAT GCC AGC GGC TCC CGG Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg ATG GCA TGG ACA CTT ATC GAG CAG CTA CAG GGC GGC AGC TAC AAG AAG Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys ATC GGC TAC TAC GAC AGC ACC AAG GAT GAT CTT TCC TGG TCC AAA ACG Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr GAC AAG TGG ATT GGA GGG TCT CCC CCA GCT GAC CAG ACC TTG GTC ATC Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile AAG ACA TTC CGT TTC CTG TCT CAG AAA CTC TTT ATC TCC GTC TCA GTT Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val CTC TCC AGC CTG GGC ATT GTT CTT GCT GTT GTC TGT CTG TCC TTT AAC Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn ATC TAC AAC TCC CAC GTT CGT TAT ATC CAG AAC TCC CAG CCC AAC CTG Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu AAC AAT CTG ACT GCT GTG GGC TGC TCA CTG GCA CTG GCT GCT GTC TTC Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe CCT CTC GGG CTG GAT GGT TAC CAC ATA GGG AGA AGC CAG TTC CCG TTT Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Ser Gln Phe Pro Phe 

GTC TGC CAG GCC CGC CTT TGG CTC TTG GGC TTG GGC TTT AGT CTG GGC

	Val	Cys	Gln	Ala	Arg	Leu	Trp	Leu	Leu	Gly	Leu	Gly	Phe	Ser	Leu	G1y	
	545					550					555	•				560	
	TAT	GGC	TCT	ATG	TTC	ACC	AAG	ATC	TGG	TGG	GTC	CAC	ACA	GTC	TTC	ACG	1728
5	Tyr	Gly	Ser	Met	Phe	Thr	Lys	Ile	Trp	Trp	Val	His	Thr	Val	Phe	Thr	
					565					570					575		
	AAG	AAG	GAG	GAG	AAG	AAG	GAG	TGG	AGG	AAG	ACC	CTA	GAG	CCC	TGG	AAA	1776
	Lys	Lys	Glu	Glu	Lys	Lys	Glu	Trp	Arg	Lys	Thr	Leu	Glu	Pro	Trp	Lys	
10				580					585					590			
	CTC	TAT	GCC	ACT	GTG	GGC	CTG	CTG	GTG	GGC	ATG	GAT	GTC	CTG	ACT	CTT	1824
•	Leu	Tyr	Ala	Thr	Val	Gly	Leu	Leu	Val	Gly	Met	Asp	Val	Leu	Thr	Leu	
			595					600					605				
. 15																	
			_												ACT		1872
	Ala		Trp	Gln	Ile	Val		Pro	Leu	His	Arg		Ile	Glu	Thr	Phe	
		610					615					620					
20															ccc		1920
		Lys	Glu	Glu	Pro		Glu	Asp	Ile	Asp		Ser	Ile	Leu	Pro		
	625					630					635					640	
	mmo	a.c	C2 C	maa	200	maa	220	220	» mc	3 3 M	3.00	mac	C/P/PI	ccc	x mm	mmc	1060
25															ATT		1968
25	Leu	GIU	.nis	Cys	5er	ser	гуs	Lys	Mec	650	THE	пр	Leu	GIŞ	Ile 655	rne	
					047					0.50					033		
	ጥልጥ	CCT	ጥልር	AAG	ccc	СТС	СТС	СТС	CTG	CTG	GGA	ልጥሮ	הלאלה	ርጥጥ	GCT	ጥልሮ	2016
															Ala		2010
30	LYL	Gry	171	660	Q <sub>1</sub>	٥٠٠	Deu		665	200	CLY			670		-1-	
50									000							•	
	GAA	ACC	AAG	AGC	GTG	TCC	АСТ	GAA	AAG	ATC	AAT	GAC	CAC	AGG	GCC	GTG	2064
															Ala		
	0_0		675					680	-4-				685	5			
35																	
	GGC	ATG	GCT	ATC	TAC	AAT	GTC	GCG	GTC	CTG	TGT	CTC	ATC	ACT	GCT	CCT	2112
															Ala		
		690			-		695				-	700					
40	GTG	ACC	ATG	ATC	СТТ	TCC	AGT	CAG	CAG	GAC	GCA	GCC	TTT	GCC	TTT	GCC	2160
-															Phe		V
	705					710				-	715				•	720	

									••				•				
	TCT	CTG	GCC	ATC	GTG	TTC	TCT	TCC	TAC	ATC	ACT	CTG	GTT	GTG	CTC	TTT	22 <del>0</del> 8
	Ser	Leu	Ala	Ile	Val	Phe	Ser	Ser	Tyr	Ile	Thr	Leu	Val	Val	Leu	Phe	
					725					730					735	•	
5	GTG	CCC	AAG	ATG	CGC	AGG	CTG	ATC	ACC	CGA	GGG	GAA	TGG	CAG	TCT	GAA	2256
	Val	Pro	Lys	Met	Arg	Arg	Leu	Ile	Thr	Arg	Gly	Glu	Trp	Gln	Ser	Glu	
				740					745					750			
	ACG	CAG	GAC	ACC	ATG	AAA	ACA	GGA	TCA	TCC	ACC	AAC	AAC	AAC	GAG	GAA	2304
10	Thr	Gln	Asp	Thr	Met	Lys	Thr	Gly	Ser	Ser	Thr	Asn	Asn	Asn	Glu	Glu	
			755					760					765				
	GAG	AAG	TCC	CGA	CTG	TTG	GAG	AAG	GAA	AAC	CGA	GAA	CTG	GAA	AAG	ATC	2352
	Glu	Lys	Ser	Arg	Leu	Leu	Glu	Lýs	Glu	Asn	Arg	Glu	Leu	Glu	Lys	Ile	
15		770					775					780					
		_															
	ATC	GCT	GAG	AAA	GAG	GAG	CGC	GTC	тст	GAA	CTG	CGC	CAT	CAG	CTC	CAG	2400
	Ile	Ala	Glu	Lys	Glu	Glu	Arg	Val	Ser	Glu	Leu	Arg	His	Gln	Leu	Gln	
	785			_		790	_				795	_				800	
20																	
	TCT	CGG	CAG	CAA	CTC	CGC	TCA	CGG	CGC	CAC	ccc	CCA	ACA	CCC	CCA	GAT	2448
	Ser	Arg	Gln	Gln	Leu	Arg	Ser	Arg	Arg	His	Pro	Pro	Thr	Pro	Pro	Asp	
		_			805			_	_	810					815	_	
25	ccc	TCT	GGG	GGC	CTT	CCC	AGG	GGA	CCC	тст	GAG	ccc	CCT	GAC	CGG	CTT	2496
	Pro	Ser	Gly	Gly	Leu	Pro	Arg	Gly	Pro	Ser	Glu	Pro	Pro	Asp	Arg	Leu	
				820					825					830			
	AGC	TGT	GAT	GGG	AGT	CGA	GTA	CAT	TTG	СТТ	TAC	AAG	TGA				2535
30	Ser	Cys	Asp	Gly	Ser	Arg	Val	His	Leu	Leu	Tyr	Lys	*				
			835					840									

(2) INFORMATION FOR SEQ ID NO: 47:

35

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 844 amino acids
  - (B) TYPE: amino acid
  - (D) TOPOLOGY: linear

40

- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:

	Met 1	Gly	Pro	Gly	Gly 5	Pro	Cys	Thr	Pro	Val 10	Gly	Trp	Pro	Leu	Pro 15	Leu
5	Leu	Leu	Val	Met 20	Ala	Ala	Gly	Val	Ala 25	Pro	Val	Trp	Ala	Ser 30	His	Ser
0	Pro	His	Leu 35	Pro	Arg	Pro	His	Pro 40	Arg	Val	Pro	Pro	His 45	Pro	Ser	Ser
	Glu	Arg 50	Arg	Ala	Val	Tyr	Ile 55	Gly	Ala	Leu	Phe	Pro 60	Met	Ser	Gly	Gly
5	Trp 65	Pro	Gly	Gly	Gln	Ala 70	Суѕ	Gln	Pro	Ala	Val 75	Glu	Met	Ala	Leu	Glu 80
	Asp	Val	Asn	Ser	Arg 85	Arg	Asp	Ile	Leu	Pro 90	Asp	Туг	Glu	Leu	Lys 95	Leu
:0	Ile	His	His	Asp 100	Ser	Lys	Cys	Asp	Pro 105	Gly	Gln	Ala	Thr	Lys 110	Tyr	Leu
25	Tyr	Glu	Leu 115	Leu	Tyr	Asn	Asp	Pro 120	Ile	Lys	Ile	Ile	Leu 125	Met	Pro	Gly
	Cys	Ser 130	Ser	Val	Ser	Thr	Leu 135	Val	Ala	Glu	Ala	Ala 140	Arg	Met	Trp	Asn
10	Leu 145	Ile	Val	Leu	Ser	Tyr 150	Gly	Ser	Ser	Ser	Pro 155	Ala	Leu	Ser	Asn	Arg 160
	Gln	Arg	Phe	Pro	Thr 165	Phe	Phe	Arg	Thr	His 170	Pro	Ser	Ala	Thr	Leu 175	His
35	Asn	Pro	Thr	Arg 180	Val	Lys	Leu	Phe	Glu 185	Lys	Trp	Gly	Trp	Lys 190	Lys	Ile
10	Ala	Thr	Ile 195	Gln	Gln	Thr	Thr	Glu 200	Val	Phe	Thr	Ser	Thr 205	Leu	Asp	Asp
	Leu	Glu	Glu	Arg	Val	Lys	Glu 215		Gly	Ile	Glu	11e	Thr	Phe	Arg	Gln

Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Thr Tyr Asp Pro Ser Ile Asn Cys Thr Val Glu Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile

Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val · 465 Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Ser Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Lys Lys Met Asn Thr Trp Leu Gly Ile Phe 

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Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Gly Ile Phe Leu Ala Tyr

Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp His Arg Ala Val

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Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu Ile Thr Ala Pro 690 695 Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala Phe Ala Phe Ala 710 715 Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu Val Val Leu Phe 730 10 Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu 740 745 Thr Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu 755 760 765 15 Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile 775 780 Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu Arg His Gln Leu Gln 790 795 Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro Pro Thr Pro Pro Asp 805 810 25 Pro Ser Gly Gly Leu Pro Arg Gly Pro Ser Glu Pro Pro Asp Arg Leu 820 825 830 Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr Lys 835 840 (2) INFORMATION FOR SEQ ID NO: 48: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2886 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 40 (ii) MOLECULE TYPE: cDNA to mRNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1..2886 10. (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48: ATG TTG CTG CTG CTA CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC 48 Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly 15 GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC 96 Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile 20 20 ATA CAC CCG CCC TGG GAA GGG GGC ATC AGG TAC CGG GGC CTG ACT CGG 144 Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg 35 40 GAC CAG GTG AAG GCT ATC AAC TTC CTG CCA GTG GAC TAT GAG ATT GAG 192 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 55 50 TAT GTG TGC CGG GGG GAG CGC GAG GTG GTG GGG CCC AAG GTC CGC AAG 240 Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys 65 70 75 TGC CTG GCC AAC GGC TCC TGG ACA GAT ATG GAC ACA CCC AGC CGC TGT 288 Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys 35 95 GTC CGA ATC TGC TCC AAG TCT TAT TTG ACC CTG GAA AAT GGG AAG GTT 336 Val Arg Ile Cys Ser Lys Ser Tyr Leu Thr Leu Glu Asn Gly Lys Val 100 105 110 40 TTC CTG ACG GGT GGG GAC CTC CCA GCT CTG GAC GGA GCC CGG GTG GAT 384 Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala Arg Val Asp 120 125 115

	ישירי	CGG	ጥርጥ	GAC	CCC	GAC	ጥፐር	САТ	CTG	GTG	GGC	AGC	TCC	CGG	AGC	ATC	432
										Val							
		130	0,70				135				3	140		5			
5		130															
J	ጥርጥ	ΔGT	CAG	GGC	CAG	TGG	AGC	ACC	CCC	AAG	CCC	CAC	TGC	CAG	GTG	ААТ	480
										Lys		_					
	145	261	<b>G1</b>	Gly	<b>U</b> 111	150	Del	****	110	2,5	155	5	<b>C</b> , 5	01	vul	160	
	147					150					133					100	
10	CCA	NCC.	CCA	CAC	תר א	CAA	ccc	CGC	CCA	GTG	ጥልር	ልጥሮ	GGG	GCA	CTC	<b>ДАДАД</b>	528
10										Val							320
	Arg	TIII	PIO	птэ		Giu	Arg	ALG	VIG	170	ıyı	***	GIY	AIG	175	ruc	
					165					170					1,73		
	ccc	አመር	NCC.	ccc	ccc	mcc.	CCA	GGG	ccc	CAG	GCC	ጥርር	CAG	ccc	ece	GTG'	576
16										Gln							3.0
15	PIO	Mec	Ser	180	GIY	ΙΙĐ	PIO	GIY	185	GIII	ALG	СУЗ	GIN	190	AIG	Val	
			•	100					103					190			
	CAC	N TOC	ccc	CTC	CAC	GAC	GTG	יית מ	AGC	CGC	) CC	GAC	איזיכי	СТС	ccc	GAC	624
							_			Arg							024
20	Giu	Mec		neu	GIU	ASD	vai	200	Ser	ALG	ALG	vsb	205	Deu	FIU	nap	
20			195					200				•	203				
	መልጥ	GAG	CTC	AAG	CTC	ልሞር	CAC	CAC	GAC	AGC	AAG	ጥርጥ	ርልጥ	CCA	GGC	C77	672
										Ser							0.2
	ıyı	210	Deu	пуs	Dea	116	215	urs	vab	Ser	בעם	220	nsp	FIO	GLY	GIII	
25		210					213					220					
25	CCC	»CC	AAC	ጥልር	СТЪ	ጥልጥ	GAG	CTG	כיזיכ	TAC	220	GAC	ССТ	ልጥሮ	AAG	ልጥሮ	720
										Tyr							
	225	1114	Буз	171	Deu	230	014	LCu	200	-3-	235	nap	110		2,3	240	
	223					250					233					240	
30	ልጥሮ	СФФ	ΑΤΥС	CCT	GGC	TGC	AGC	тст	GTC	TCC	ACG	CTG	GTG	GCT	GAG	GCT	768
30			•							Ser							,
	116	Deu	1160	110	245	C, C	001	502	***	250	****	200	741		255		
					2.5												
	CCT	AGG	АТС	тсс	AAC	СТС	АТТ	GTG	СТТ	TCC	тат	GGC	TCC	AGC	TCA	CCA	816
35										Ser							
33	niu	na g		260					265		-3-	,		270			
				200													
	GCC	СТС	тса	AAC	CGG	CAG	CGT	TTC	ccc	ACT	TTC	ттс	CGA	ACG	CAC	CCA	864
										Thr							
40	ALG	ے۔ د	275		••• 9		3	280					285				
₩.			213										200				ν
	ጥሮኔ	GCC	202	ርጥር	CAC	AAC	ССТ	ACC	CGC	GTG	AAA	כתר	ጥጥጥ	GAÁ	AAG	TGG	912
										Val							
	ser	VIG		חבת	****	£1911			9	741	-7.9	J-cu	1116	JIU	_, _		

	H18	65-1	wo	SEQ				4	40								
	CAG	GAG	GCA	CCG	CTG	GCC	ТАТ	GAT	GCC	ATC	TGG	GCC	TTG	GCA	CTG	GCC.	1440
		450					455					460			•		
40	Lys		Thr	Lys	Arg	Leu	-	Arg	His	Pro	Glu		Thr	Gly	Gly	Phe	v
										CCT							1392
			433					71 18 U					447				
	Ala	Asn	Thr 435	Arg	ser	TTE	ser	Asn 440	met	Thr	ser	GIN	G1u	rne	vai	GIU	
35										ACA		_	_		_	_	1344
		_	_			. –	_ =										
				420					425					430			
										Glu							
50	GAG	GCG	GTG	GAG	GGC	CAC	ATC	ACA	ACT	GAG	ATT	GTC	ATG	CTG	AAT	ССТ	1296
30					403					#T0					413		
	Phe	Lys	Ile	Tyr	Asp 405	Pro	Ser	Ile	Asn	Cys 410	Thr	Val	Asp	Glu	Met 415	Thr	
										TGC							1248
25	385					390					395					400	
	Gly	Lys	Lys	Tyr	Val	Trp	Phe	Leu	Ile	Gly	Trp	Tyr	Ala	Asp	Asn	Trp	
	GGG	AAG	AAG	TAC	GTC	TGG	TTC	CTC	ATT	GGG	TGG	TAT	GCT	GAC	AAT	TGG	1200
		310					373					300					
20	rnr	370	ALA	Arg	ьys	vaı	375	Cys	GIU	Val	тух	380	GIÜ	Arg	ьец	rne	
••						_				GTG							1152
																	_
			355					360					365				
-										Ile							
15	AAC	CTG	AAG	CGC	CAG	GAT	GCC	CGA	ATC	ATC	GTG	GGA	CTT	TTC	ТАТ	GAG	1104
				340					345					350			
	Ile	Thr	Phe	_	Gln	Ser	Phe	Phe		Asp	Pro	Ala	Val		Val	Lys	
										GAT							1056
10																	
				_	325					330				_	335		
										Val							
	TCG	ACT	CTG	GAC	GAC	CTG	GAG	GAA	CGA	GTG	AAG	GAG	GCT	GGA	ATT	GAG	1008
5	305					310					315					320	
	Gly	Trp	Lys	Lys	Ile		Thr	Ile	Gln	Gln		Thr	Glu	Val	Phe		
	GGC	TGG	AAG	AAG	ATT	GCT	ACC	ATC	CAG	CAG	ACC	ACT	GAG	GTC	TTC	ACT	960
		290					295					300					
									-				•				

	Gln	Glu	Ala	Pro	Leu	Ala	Tyr	Asp	Ala	Ile	Trp	Ala	Leu	Ala	Leu	Ala	· <del></del>
	465					470					475					480	
	CTG	AAC	AAG	ACA	тст	GGA	GGA	GGC	GGC	CGT	TCT	GGT	GTG	CGC	CTG	GAG	1488
5	Leu	Asn	Lys	Thr	Ser	Gly	Gly	Gly	Gly	Arg	Ser	Gly	Val	Arg	Leu	Glu	
					485					490					495		
	GAC	TTC	AAC	TAC	AAC	AAC	CAG	ACC	ATT	ACC	GAC	CAA	ATC	TAC	CGG	GCA	1536
	Asp	Phe	Asn	Tyr	Asn	Asn	Gln	Thr	Ile	Thr	Asp	Gln	Ile	Tyr	Arg	Ala	
10				500					505					510			
	ATG	AAC.	TCT	TCG	TCC	TTT	GAG	GGT	GTC	TCT	GGC	CAT	GTG	GTG	TTT	GAT	1584
	Met	Asn	Ser	Ser	Ser	Phe	Glu	Gly	Val	Ser	Gly	His	Val	Val	Phe	Asp	
			515					520					525				
15																	
	GCC	AGC	GGC	TCT	CGG	ATG	GCA	TGG	ACG	CTT	ATC	GAG	CAG	СТТ	CAG	GGT	1632
	Ala	Ser	Gly	Ser	Arg	Met	Ala	Trp	Thr	Leu	Ile	Glu	Gln	Leu	Gln	Gly .	
		530	•				535					540					
20	GGC	AGC	TAC	AAG	AAG	ATT	GGC	TAC	TAT	GAC	AGC	ACC	AAG	GAT	GAT	CTT	1680
	Gly	Ser	Tyr	Lys	Lys	Ile	Gly	Tyr	Tyr	Asp	Ser	Thr	Lys	Asp	Asp	Leu	
	545					550					555					560	
	TCC	TGG	TCC	AAA	ACA	GAT	AAA	TGG	ATT	GGA	GGG	TCC	CCC	CCA	GCT	GAC	1728
25	Ser	Trp	Ser	Lys	Thr	Asp	Lys	Trp	Ile	Gly	Gly	Ser	Pro	Pro	Ala	Asp	
					565					570					575		
	CAG	ACC	CTG	GTC	ATC	AAG	ACA	TTC	CGC	TTC	CTG	TCA	CAG	AAA	CTC	TTT	1776
	Gln	Thr	Leu	Val	Ile	Lys	Thr	Phe	Arg	Phe	Leu	Ser	Gln	Lys	Leu	Phe	
30				580					585					590			
	ATC	TCC	GTC	TCA	GTT	CTC	TCC	AGC	CTG	GGC	ATT	GTC	CTA	GCT	GTT	GTC	1824
	Ile	Ser	Val	Ser	Val	Leu	Ser	Ser	Leu	Gly	Ile	Val	Leu	Ala	Val	Val	
			595					600					605				
35																	
	TGT	CTG	TCC	TTT	AAC	ATC	TAC	AAC	TCA	CAT	GTC	CGT	TAT	ATC	CAG	AAC	1872
	Суѕ	Leu	Ser	Phe	Asn	Ile	Tyr	Asn	Ser	His	Val	Arg	Tyr	Ile	Gln	Asn	
		610					615					620					
40	TCA	CAG	CCC	AAC	CTG	AAC	AAC	CTG	ACT	GCT	GTG	GGC	TGC	TCA	CTG	GCT	1920ս
	Ser	Gln	Pro	Asn	Leu	Asn	Asn	Leu	Thr	Ala	Val	Gly	Суѕ	Ser	Leu	Ala	
	625					630					635					640	

								•	-				•					
	TTA	GCT	GCT	GTC	TTC	ccc	CTG	GGG	CTC	GAT	GGT	TAC	CÁC	ATT	GGG	AGG	••	19 <del>6</del> 8
	Leu	Ala	Ala	Val	Phe	Pro	Leu	Gly	Leu	Asp	Gly	Tyr	His	Ile	Gly	Arg		
					645					650					655			
5	AAC	CAG	TTT	CCT	TTC	GTC	TGC	CAG	GCC	CGC	CTC	TGG	CTC	CTG	GGC	CTG		2016
	Asn	Gln	Phe	Pro	Phe	Val	Cys	Gln	Ala	Arg	Leu	Trp	Leu	Leu	Gly	Leu		
			•	660					665					670			•	
	GGC	TTT	AGT	CTG	GGC	TAC	GGT	TCC	ATG	TTC	ACC	AAG	ATT	TGG	TGG	GTC	•	2064
10	Gly	Phe	Ser	Leu	Gly	Tyr	Gly	Ser	Met	Phe	Thr	Lys	Ile	Trp	Trp	Val		
			675					680					685					
	CAC	ACG	GTC	TTC	ACA	AAG	AAG	GAA	GAA	AAG	AAG	GAG	TGG	AGG	AAG	ACT		2112
	His	Thr	Val	Phe	Thr	Lys	Lys	Glu	Glu	Lys	Lys	Glu	Trp	Arg	Lys	Thr		
15		690					695					700						
	CTG	GAA	CCC	TGG	AAG	CTG	TAT	GCC	ACA	GTG	GGC	CTG	CTG	GTG	GGC	ATG		2160
	Leu	Glu	Pro	Trp	Lys	Leu	Tyr	Ala	Thr	Val	Gly	Leu	Leu	Val	Gly	Met		
	705					710					715					720		
20																		
	GAT	GTC	CTC	ACT	CTC	GCC	ATC	TGG	CAG	ATC	GTG	GAC	CCT	CTG	CAC	CGG		2208
	Asp	Val	Leu	Thr	Leu	Ala	Ile	Trp	Gln	Ile	Val	Asp	Pro	Leu	His	Arg		•
					725					730					735			
•																		
25	ACC	ATT	GAG	ACA	TTT	GCC	AAG	GAG	GAA	CCT	AAG	GAA	GAT	ATT	GAC	GTC		2256
	Thr	Ile	Glu	Thr	Phe	Ala	Lys	Glu	Glu	Pro	Lys	Glu	Asp	Ile	Asp	Val		
				740					745					750				
											TCC							2304
30	Ser	Ile		Pro	Gln	Leu	Glu		Cys	Ser	Ser	Arg		Met	Asn	Thr		
			755					760					765					
							-				CTG							2352
	Trp		GIÄ	Ile	Phe	Tyr	_	Tyr	Lys	Gly	Leu		Leu	Leu	Leu	Gly	•	
35		770					775					780						
	1.00	mma	omm.	00m	m> m	63.6			3 CM	cmc	maa		~~~		1 mg			2400
											TCC							2400
		Pne	ren	Ala	ıyr		Thr	гÀг	ser	vaı	Ser	rnr	GIU	rys	ше			
4-5	785					790					795					800		
40	a	<b></b>					<b>1</b>			m		050	ac:		or->	mo c		D 4 4 0
											AAT							2448
	Asp	His	Arg	Ala		Gly	Met	Ala	ITe		Asn	val	Ala	val		Cys		
					805					810					815			

																•	
	CTC	ATC	ACT	GCT	ССТ	GTC	ACC	ATG	ATT	CTG	TCC	AGC	CAG	CAG	GAT	GCA	2496
	Leu	Ile	Thr	Ala	Pro	Val	Thr	Met	Ile	Leu	Ser	Ser	Gln	Gln	Asp	Ala	
				820					825					830			
5																	
	GCC	TTT	GCC	TTT	GCC	TCT	CTT	GCC	ATA	GTT	TTC	TCC	TCC	TAT	ATC	ACT	2544
	Ala	Phe	Ala	Phe	Ala	Ser	Leu	Ala	Ile	Val	Phe	Ser	Ser	Tyr	Ile	Thr	
			835					840					845				
10	CTT	GTT	GTG	CTC	TTT	GTG	CCC	AAG	ATG	CGC	AGG	CTG	ATC	ACC	CGA	GGG	2592
	Leu	Val	Val	Leu	Phe	Val	Pro	Lys	Met	Arg	Arg	Leu	Ile	Thr	Arg	Gly	
		850					855					860					
												ACA					2640
15		Trp	Gln	Ser	Glu		Gln	Asp	Thr	Met	-	Thr	Gly	Ser	Ser	Thr	
	865					870					875					880	
					_							GAG					2688
••	Asn	Asn	Asn	GIU		Glu	гЛS	Ser	Arg		Leu	Glu	Lys	GIu		Arg	
20					885					890					895		
	CAA	CTC	CAA	N N C	አጥሮ	אייניי א	CCT	CAC	***	CAC	CAC	CGT	CTTC	TO TO	CAA	СТС	2736
												Arg					2730
	014		014	900				014	905	014	014	9	<b>,</b>	910	014	Deu	
25				,,,,					,,,					320			
	CGC	CAT	CAA	CTC	CAG	TCT	CGG	CAG	CAG	CTC	CGC	TCC	CGG	CGC	CAC	CCA	2784
	Arg	His	Gln	Leu	Gln	Ser	Arg	Gln	Gln	Leu	Arg	Ser	Arg	Arg	His	Pro	
	_		915				_	920			-		925	_			
30	CCG	ACA	CCC	CCA	GAA	CCC	TCT	GGG	GGC	CTG	CCC	AGG	GGA	CCC	ССТ	GAG	2832
	Pro	Thr	Pro	Pro	Glu	Pro	Ser	Gly	Gly	Leu	Pro	Arg	Gly	Pro	Pro	Glu	
		930					935					940					
	CCC	CCC	GAC	CGG	CTT	AGC	TGT	GAT	GGG	AGT	CGA	GTG	CAT	TTG	CTT	TAT	2880
35	Pro	Pro	Asp	Arg	Leu	Ser	Cys	Asp	Gly	Ser	Arg	Val	His	Leu	Leu	Tyr	
	945					950					955					960	
	AAG	TGA															2886
	Lys	*	•													•	
40																	1.

(2) INFORMATION FOR SEQ ID NO: 49:

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- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 961 amino acids
  - (B) TYPE: amino acid
- 5 (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: protein
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 49:

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly

1 . 5 . 10 . 15

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile
15 20 25 30

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg
35 40 45

20 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 55 60

Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys
65 70 75 80

Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys
85 90 95

Val Arg Ile Cys Ser Lys Ser Tyr Leu Thr Leu Glu Asn Gly Lys Val

100 105 110

Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala Arg Val Asp 115 120 125

Phe Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser Arg Ser Ile 130 135 140

Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys Gln Val Asn 145 150 155 160

Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe 165 170 175

Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile Leu Met Pro Gly Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn Leu Ile Val Leu Ser Tyr Gly Ser Ser Pro Ala Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser Ala Thr Leu His Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly Trp Lys Lys Ile Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr Ser Thr Leu Asp Asp Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile Thr Phe Arg Gln Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Ile Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr

PCT/SE98/01947

WO 99/21890 Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala 15 Leu Asn Lys Thr Ser Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn 

Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala

	Leu	Ala	Ala	Val	Phe 645	Pro	Leu	Gly	Leu	Asp 650	Gly	Tyr	His	Ile	Gly 655	Arg
5	Asn	Gln	Phe	Pro 660	Phe	Val	Cys	Gln	Ala 665	Arg	Leu	Trp	Leu	Leu 670	Gly	Leu
10	Gly	Phe	Ser 675	Leu	Gly	Tyr	Gly	Ser 680	Met	Phe	Thr	Lys	Ile 685	Trp	Trp	Val
	His	Thr 690	Val	Phe	Thr	Lys	Lys 695	Glu	Glu	Lys	Lys	Glu 700	Trp	Arg	Lys	Thr
15	Leu 705	Glu	Pro	Trp	Lys	Leu 710	Tyr	Ala	Thr	Val	Gly 715	Leu	Leu	Val	Gly	Met 720
	Asp	Val	Leu	Thr	Leu 725	Ala	Ile	Trp	Gln	Ile 730	Val	Asp	Pro	Leu	His 735	Arg
20	Thr	Ile	Glu	Thr 740	Phe	Ala	Lys	Glu	Glu 745	Pro	Lys	Glu	Asp	Ile 750	Asp	Val
25	Ser	Ile	Leu 755	Pro	Gln	Leu	Glu	His 760	Суз	Ser	Ser	Arg	Lys 765	Met	Asn	Thr
	Trp	Leu 770	Gly	Ile	Phe	Tyr	Gly 775	Tyr	Lys	Gly	Leu	Leu 780	Leu	Leu	Leu	Gly
30	Ile 785	Phe	Leu	Ala	Tyr	Glu 790	Thr	Lys	Ser	Val	Ser 795	Thr	Glu	Lys	Ile	Asn 800
	Asp	His	Arg	Ala	Val 805	Gly	Met	Àla	Ile	Туг 810	Asn	Val	Ala	Val	Leu 815	Суз
35	Leu	Ile	Thr	Ala 820	Pro	Val	Thr	Met	Ile 825	Leu	Ser	Ser	Gln	Gln 830	Asp	Ala
40	Ala	Phe	Ala 835	Phe	Ala	Ser	Leu	Ala 840	Ile	Val	Phe	Ser	Ser 845	Tyr	Ile	Thr
	Leu	Val 850	Val	Leu	Phe	Val	Pro 855	Lys	Met	Arg	Arg	Leu 860	Ile	Thr	Arg	Gly

Glu Trp Gln Ser Glu Ala Gln Asp Thr Met Lys Thr Gly Ser Ser Thr 865 870 875 880

Asn Asn Asn Glu Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg 885 890 895

Glu Leu Glu Lys Ile Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu 900 905 910

10 Arg His Gln Leu Gln Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro 915 920 925

Pro Thr Pro Pro Glu Pro Ser Gly Gly Leu Pro Arg Gly Pro Pro Glu 930 935 940

Pro Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr 945 950 955 960

Lys

20

15

- (2) INFORMATION FOR SEQ ID NO: 50:
- 25 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2535 base pairs

(B) TYPE: nucleic acid(C) STRANDEDNESS: single(D) TOPOLOGY: linear

30

- (ii) MOLECULE TYPE: cDNA to rRNA
- (iii) HYPOTHETICAL: NO
- 35 (iv) ANTI-SENSE: NO
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo sapiens
- 40 (ix) FEATURE:
  - (A) NAME/KEY: CDS
  - (B) LOCATION: 1..2535

		(xi)	SEC	JUENO	CE DE	ESCRI	PTIC	on: 9	SEO .	D NO	): 50	):	•	•	•			
		\^_/	Ų DĄ	202					. g			•						
	ATG	GGG	ccc	GGG	GCC	CCT	TTT	GCC	CGG	GTG	GGG	TGG	CCA	CTG	CCG	CTT		48
	Met	Gly	Pro	Gly	Ala	Pro	Phe	Ala	Arg	Val	Gly	Trp	Pro	Leu	Pro	Leu		
5	1				5					10					15			
					GCG													96
	Leu	Val	Val		Ala	Ala	Gly	Val		Pro	Val	Trp	Ala		His	Ser		
				20					25					30				
10	ccc	<sub>С</sub> ът	רתכ	CCG	CGG	ССТ	CAC	ጥርር	CGG	GTC	CCC	CCG	CAC	CCC	ጥርር	ጥሮል		144
					Arg								_					
			35					40					45					
15	GAA	CGG	CGC	GCA	GTG	TAC	ATC	GGG	GCA	CTG	TTT	CCC	ATG	AGC	GGG	GGC		192
	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	Ala	Leu	Phe	Pro	Met	Ser	Gly	Gly		
		50					55					60						
					CAG													240
20	•	Pro	СТĀ	GIY	Gln	70	cys	GIN	PIO	AIA	75	GIU	met	ATA	ren	80 80		
	65					70					, ,					80		
	GAC	GTG	AAT	AGC	CGC	AGG	GAC	ATC	CTG	CCG	GAC	TAT	GAG	CTC	AAG	CTC		288
	Asp	Val	Asn	Ser	Arg	Arg	Asp	Ile	Leu	Pro	Asp	Tyr	Glu	Leu	Lys	Leu		
25					85					90					95			
																	·	
					AGC													336
	Ile	His	His	_	Ser	Lys	Cys	Asp		Gly	Gln	Ala	Thr		Tyr	Leu		
				100					105					110				
30	ייימייי	GNG	CTG	CTC	TAC	ልልሮ	GAC	ССТ	እጥር	AAG	ልጥሮ	ልጥሮ	ርጥጥ	ΔΤΥС	CCT	GGC		384
					Tyr													301
	-1-		115				2	120					125					
35	TGC	AGC	TCT	GTC	TCC	ACG	CTG	GTG	GCT	GAG	GCT	GCT	AGG	ATG	TGG	AAC		432
	Cys	Ser	Ser	Val	Ser	Thr	Leu	Val	Ala	Glu	Ala	Ala	Arg	Met	Trp	Asn		
		130					135					140						
																CGG	•	480
40		Ile	Val	Leu	Ser	_	Gly	Ser	Ser	Ser		Ala	Leu	Ser	Asn			υ
	145					150					155				•	160		
	ርልር	ርርጥ	ጥጥር	CCC	ACT	ጥጥር	ጥጥ	CGA	ACG	CAC	CCA	TCA	GCC	ACA	CTC	CAC		528
	CAG																	•
	H18	65-1	wo	SEO					49									<del></del>

									•				•				
	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	His	Pro	Ser	Ala	Thr	Leu	His	
					165					170					175		
	AAC	CCT	ACC	CGC	GTG	AAA	CTC	TTT	GAA	AAG	TGG	GGC	TGG	AAG	AAG	ATT	576
5	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe	Glu	Lys	Trp	Gly	Trp	Lys	Lys	Ile	
				180					185					190			
	GCT	ACC	ATC	CAG	CAG	ACC	ACT	GAG	GTC	TTC	ACT	TCG	ACT	CTG	GAC	GAC	624
	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	Val	Phe	Thr	Ser	Thr	Leu	Asp	Asp	
10			195					200					205				
	CTG	GAG	GAA	CGA	GTG	AAG	GAG	GCT	GGA	ATT	GAG	ATT	ACT	TTC	CGC	CAG	672
	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala	Gly	Ile	Glu	Ile	Thr	Phe	Arg	Gln	
		210					215					220					
15																	
	AGT	TTC	TTC	TCA	GAT	CCA	GCT	GTG	ccc	GTC	AAA	AAC	CTG	AAG	CGC	CAG	720
	Ser	Phe	Phe	Ser	Asp	Pro	Ala	Val	Pro	Val	Lys	Asn	Leu	Lys	Arg	Gln	
	225					230					235					240	
20	GAT	GCC	CGA	ATC	ATC	GTG	GGA	CTT	TTC	TAT	GAG	ACT	GAA	GCC	CGG	AAA	768
	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	Phe	Tyr	Glu	Thr	Glu	Ala	Arg	Lys	
					245					250					255		
	GTT	ттт	TGT	GAG	GTG	TAC	AAG	GAG	CGT	CTC	TTT	GGG	AAG	AAG	TAC	GTC	816
25	Val	Phe	Cys	Glu	Val	Tyr	Lys	Glu	Arg	Leu	Phe	Gly	Lys	Lys	Tyr	Val	
				260					265					270			
	TGG	TTC	CTC	ATT	GGG	TGG	TAT	GCT	GAC	AAT	TGG	TTC	AAG	ATC	TAC	GAC	864
	Trp	Phe	Leu	Ile	Gly	Trp	Tyr	Ala	Asp	Asn	Trp	Phe	Lys	Ile	Tyr	Asp	
30			275					280					285				
	CCT	TCT	ATC	AAC	TGC	ACA	GTG	GAT	GAG	ATG	ACT	GAG	GCG	GTG	GAG	GGC	912
	Pro	Ser	Ile	Asn	Суз	Thr	Val	Asp	Glu	Met	Thr	Glu	Ala	Val	Glu	Gly	
		290					295					300					
35																	
	CAC	ATC	ACA	ACT	GAG	ATT	GTC	ATG	CTG	AAT	CCT	GCC	AAT	ACC	CGC	AGC	960
	His	Ile	Thr	Thr	Glu	Ile	Val	Met	Leu	Asn	Pro	Ala	Asn	Thr	Arg	Ser	
	305					310					315					320	
40	ATT	TCC	AAC	ATG	ACA	TCC	CAG	GAA	TTT	GTG	GAG	AAA	CTA	ACC	AAG	CGA	1008
	Ile	Ser	Asn	Met	Thr	Ser	Gln	Glu	Phe	Val	Glu	Lys	Leu	Thr	Lys	Arg	,
					325					330		•			^335		

				500					505					510			
	Ile	Tyr	Asn		His	Val	Arg	Tyr		Gln	Asn	Ser	Gln		Asn	Leu	
			AAC												_		1536
40			•														υ
					485					490					495		
	Leu	Ser	Ser	Leu	Gly	Ile	Val	Leu	Ala	Val	Val	Cys	Leu	Ser	Phe	Asn	
	CTC	TCC	AGC	CTG	GGC	ATT	GTC	СТА	GCT	GTT	GTC	TGT	CTG	TCC	TTT	AAC	1488
35	400					41/0					4/3					40V	
16	Lys 465	Thr	Phe	Arg	rne	Leu 470	ser	GIN	гÀз	ьeu	Phe 475	тте	ser	val	ser	Val 480	
			TTC														1440
		450					455					460					
30	Asp	Lys	Trp	Ile	Gly	Gly	Ser	Pro	Pro	Ala	Asp	Gln	Thr	Leu	Val	Ile	
	GAT	AAA	TGG	ATT	GGA	GGG	TCC	ccc	CCA	GCT	GAC	CAG	ACC	CTG	GTC	ATC	1392
			435					440					445				
	Ile	Gly	Tyr	Tyr	Asp	Ser	Thr	_	Asp	Asp	Leu	Ser	_	Ser	Lys	Thr	
25			TAC														1344
									_	_							
v				420					425					430			
	Met	Ala	Trp	Thr	Leu	Ile	Glu	Gln	Leu	Gln	Gly	Gly	Ser	Tyr	Lys	Lys	
	ATG	GCA	TGG	ACG	CTT	ATC	GAG	CAG	CTT	CAG	GGT	GGC	AGC	TAC	AAG	AAG	1296
20															717		
	rne	GIU	GTÀ	val	405	GIĀ	uis	val	val	410	ьгр	wig	ser	стλ	415	VIA	
			GGT Gly														1248
		<b></b>	00=	ama	mc=	000	03 m	OE-C	OFF	mere.	<b>01</b> T	000	1.00	000	me=	999	1040
15	385					390					395					400	
	Asn	Gln	Thr	Ile	Thr	Asp	Gln	Ile	Tyr	Arg	Ala	Met	Asn	Ser	Ser	Ser	
	AAC	CAG	ACC	ATT	ACC	GAC	CAA	ATC	TAC	CGG	GCA	ATG	AAC	TCT	TCG	TCC	1200
	3	370	- 3	- 3	2	-	375	_	-		_ ==	380			- <b></b>	v- <del>-</del>	
10			Gly														
	GGA	GGA	GGC	GGC	CGT	TCT	GGT	GTG	CGC	CTG	GAG	GAC	TTC	AAC	TAC	AAC	1152
			355					360					365				
	Ala	Tyr	Asp	Ala	Ile	Trp	Ala		Ala	Leu	Ala	Leu		Lys	Thr	Ser	
5			GAT														1104
				340					345					350			
			Arg														
	CTYC	ΔΔΔ	AGA	CAC	CCT	GAG	GAG	ACA	GGA	GGC	TTC	CAG	GÁG	GCÁ	CCG	CTG	10 <del>5</del> 6

												٠	•				
	AAC	AAC	CTG	ACT	GCT	GTG	GGC	TGC	TCA	CTG	GCT	TTA	GCT	GCT	GTC	TTC	1584
					Ala												
			515					520					525				
			313					320					223				
5	000	ama	000	CMC	GAT	CCM	mac.	CAC	N TOTO	ccc	NCC.	220	C2.C	mmm	CCM	mmC	1632
									_								1032
	Pro		GIY	Leu	Asp	GIŞ	_	nıs	116	GIY	Arg		GIN	Pne	Pro	Pne	
		530					535					540					
10					CGC												1680
	Val	Cys	Gln	Ala	Arg	Leu	Trp	Leu	Leu	Gly		Gly	Phe	Ser	Leu	Gly	
	545					550					555					560	
	TAC	GGT	TCC	ATG	TTC	ACC	AAG	ATT	TGG	TGG	GTC	CAC	ACG	GTC	TTC	ACA	1728
15	Tyr	Gly	Ser	Met	Phe	Thr	Lys	Ile	Trp	Trp	Val	His	Thr	Val	Phe	Thr	
					565					570					575		
	AAG	AAG	GAA	GAA	AAG	AAG	GAG	TGG	AGG	AAG	ACT	CTG	GAA	CCC	TGG	AAG	1776
	Lys	Lys	Glu	Glu	Lys	Lys	Glu	Trp	Arg	Lys	Thr	Leu	Glu	Pro	Trp	Lys	
20				580					585					590			
	CTG	TAT	GCC	ACA	GTG	GGC	CTG	CTG	GTG	GGC	ATG	GAT	GTC	CTC	ACT	CTC	1824
	Leu	Tyr	Ala	Thr	Val	Gly	Leu	Leu	Val	Gly	Met	Asp	Val	Leu	Thr	Leu	
			595					600					605				•
25																	
	GCC	ATC	TGG	CAG	ATC	GTG	GAC	CCT	CTG	CAC	CGG	ACC	АТТ	GAG	ACA	TTT	1872
	Ala	Ile	Trp	Gln	Ile	Val	Asp	Pro	Leu	His	Arg	Thr	Ile	Glu	Thr	Phe	
		610					615					620					
			•														
30	GCC	AAG	GAG	GAA	CCT	AAG	GAA	GAT	ATT	GAC	GTC	TCT	ATT	CTG	CCC	CAG	1920
	Ala	Lys	Glu	Glu	Pro	Lys	Glu	Asp	Ile	Asp	Val	Ser	Ile	Leu	Pro	Gln	
	625					630					635					640	
	CTG	GAG	CAT	TGC	AGC	TCC	AGG	AAG	ATG	AAT	ACA	TGG	CTT	GGC	ATT	TTC	1968
35	Leu	Glu	His	Cvs	Ser	Ser	Arq	Lvs	Met	Asn	Thr	Trp	Leu	Gly	Ile	Phe	
•••					645		•	-		650	•	_		_	655		
	ጥልጥ	GGT	TAC	AAG	GGG	CTG	CTG	CTG	CTG	CTG	GGA	ATC	TTC	CTT	GCT	TAT	2016
					Gly												
40	* Y *	UL y	* Y *	660					665		1			670		_	
40				000					000					J.J			υ
	030	200	220	N C/m	GTG	TICC.	אכייי	GAC	A A C	ልጥሮ	ልልጥ	САТ	CAC	CGG	ᢅᢎᠸ᠇᠇	GTG	2064
																	2004
	GIU	Thr	гÃг	ser	Val	ser	Inr	GIU	гÀг	TIG	ASII	Asp	urs	wrâ	VIG	AGT	
								_									

			675					680					685			•	-
															GCT		2112
5	Gly	Met 690	Ala	Ile	Tyr	Asn	Val 695	Ala	Val	Leu	Cys	Teu 700	Ile	Thr	Ala	Pro	
															TTT Phe		2160
10	705	1111	Mec	116	Бец	710	Ser	GIN	GIII	ASP	715	ATG	rne	AIG	rne	720	
10															CTC Leu		2208
	361	Dea	ALU	110	725	1110	501	561	-7-	730		200	,,,	V	735	• •••	
15															TCG Ser		2256
	Vai	110	БYЗ	740	mrg	1119	Deu	110	745	9	CLI	Ciu		750	001	024	
20				-											GAG Glu		2304
20			755			-1-		760					765				
												_		_	AAG Lys	_	2352
25		770					775	-			_	780			-		
															CTC Leu		2400
30	785			-		790					795					800	
															CCA Pro		2448
		•			805					810					815		
35															CGG Arg		2496
			-	820					825					830			
40				GGG Gly									TGA *				2535
-		-	835	_		_		840									

(2) INFORMATION FOR SEQ ID NO: 51:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 844 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 51:

Met Gly Pro Gly Ala Pro Phe Ala Arg Val Gly Trp Pro Leu Pro Leu

1 5 10 15

Leu Val Val Met Ala Ala Gly Val Ala Pro Val Trp Ala Ser His Ser

20 25 30

Pro His Leu Pro Arg Pro His Ser Arg Val Pro Pro His Pro Ser Ser 35 40 45

20

Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe Pro Met Ser Gly Gly 50 55 60

Trp Pro Gly Gln Ala Cys Gln Pro Ala Val Glu Met Ala Leu Glu
25 65 70 75 80

Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu 85 90 95

30 Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala Thr Lys Tyr Leu 100 105 110

Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile Leu Met Pro Gly
115 120 125

35

Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn 130 135 140

Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala Leu Ser Asn Arg
40 145 150 155 160

Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser Ala Thr Leu His
165 170 175

	Asn	Pro	Thr	Arg 180	Val	Lys	Leu	Phe	Glu 185	Lys	Trp	Gly	Trp	Lys 190	Lys	Ile	
5	Ala	Thr	Ile 195	Gln	Gln	Thr	Thr	Glu 200	Val	Phe	Thr	Ser	Thr 205	Leu	Asp	Asp	
10	Leu	Glu 210	Glu	Arg	Val	Lys	Glu 215	Ala	Gly	Ile	Glu	Ile 220	Thr	Phe	Arg	Gln	
	Ser 225	Phe	Phe	Ser	Asp	Pro 230	Ala	Val	Pro	Val	Lys 235	Asn	Leu	Lys	Arg	Gln 240	
15	Asp	Ala	Arg	Ile	Ile 245	Val	Gly	Leu	Phe	Туг 250	Glu	Thr	Glu	Ala	Arg 255	Lys	
	Val	Phe	Cys	Glu 260	Val	Tyr	Lys	Glu	Arg 265	Leu	Phe	Gly	Lys	Lys 270	Tyr	Val	
20	Trp	Phe	Leu 275	Ile	Gly	Trp	Tyr	Ala 280	Asp	Asn	Trp	Phe	Lys 285	Ile	Tyr	Asp	
25	Pro	Ser 290	Ile	Asn	Cys	Thr	Val 295	Asp	Glu	Met	Thr	Glu 300	Ala	Val	Glu	Gly	
	His 305	Ile	Thr	Thr	Glu	Ile 310	Val	Met	Leu	Asn	Pro 315	Ala	Asn	Thr	Arg	Ser 320	
30	Ile	Ser	Asn	Met	Thr 325	Ser	Gln	Glu	Phe	Val 330	Glu	Lys	Leu	Thr	Lys 335	Arg	
	Leu	Lys	Arg	His 340	Pro	Glu	Glu		Gly 345	-	Phe	Gln	Glu	Ala 350	Pro	Leu	
35	Ala	Tyr	Asp 355	Ala	Ile	Trp	Ala	Leu 360	Ala	Leu	Ala	Leu	Asn 365	Lys	Thr	Ser	
40	Gly	Gly 370	Gly	Gly	Arg	Ser	Gly 375	Val	Arg	Leu	Glu	Asp 380	Phe	Asn	Tyr	Asn	
	Asn 385	Gln	Thr	Ile	Thr	Asp 390	Gln	Ile	Tyr	Arg	Ala 395	Met	Asn	Ser	Ser	Ser 400	

Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Asn Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln

625 630 635 640

Leu Glu His Cys Ser Ser Arg Lys Met Asn Thr Trp Leu Gly Ile Phe  $645 \hspace{1.5cm} 650 \hspace{1.5cm} 655 \hspace{1.5cm}$ 

Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Gly Ile Phe Leu Ala Tyr
660 665 670

Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp His Arg Ala Val
10 675 680 685

Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu Ile Thr Ala Pro 690 695 700

Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala Phe Ala Phe Ala 705 710 715 720

Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu Val Val Leu Phe
725 730 735

20

Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu
740 745 750

Ala Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu
25 755 760 765

Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile 770 775 780

30 Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu Arg His Gln Leu Gln 785 790 795 800

Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro Pro Thr Pro Pro Glu 805 810 815

35

40

Pro Ser Gly Gly Leu Pro Arg Gly Pro Pro Glu Pro Pro Asp Arg Leu 820 825 830

Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr Lys 835 840

(2) INFORMATION FOR SEQ ID NO: 52:

		(i)	(1	QUENC A) LI 3) TY	engti (PE :	1: 2: nuc	395 l leic	oase acid	pai:	rs							
,			-	) TO				_	316								
		(ii)	MOI	LECUI	LE T	PE:	cDN	A to	mRN	Ą							
10	(	(iii)	HY:	РОТНІ	ETIC!	AL: 1	NO										
		(iv)	AN'	ri-si	ENSE	: NO											
		(vi		IGIN?													
15			(2	A) . OI	RGAN:	ISM:	Can	is fa	amil:	iari	3						
		(ix)		ATURI		27772	ana										
			•	A) NA B) L(				895									
20		(vi	SE(	QUENC	ות פי	SCR	<b>የ</b> የተመሰ	N - 9	SEO 1	ות חז	o 51	2.					
		(***	, 52,	201111		JOC11.			Jug .		. J.	••					
				CTG Leu													48
25	1				5					10					15		
	ccc	CCG	GGC	GCG	GGC	GGG	GCA	CAG	ACC	ccc	AAC	GCC	ACC	TCG	GAA	GGT	96
	Pro	Pro	Gly	Ala 20	Gly	Gly	Ala	Gln	Thr 25	Pro	Asn	Ala	Thr	Ser 30	Glu	Gly	
30																	
				ATA Ile													144
	-2-		35					40					45		2		
35	CTG	ACT	CGT	GAC	CAG	GTG	AAG	GCT	ATC	AAC	TTC	CTG	CCG	GTG	GAC	TAT	192
	Leu		Arg	Asp	Gln	Val	_	Ala	Ile	Asn	Phe		Pro	Val	Asp	Tyr	
		50					55					60					
			•	TAT													240
40	Glu 65	Ile	Glu	Tyr	Val	70	Arg	Gly	Glu	Arg	G1u 75	Val	Val	Gly	Pro	Lys 80	
															•		
	GTC	CGA	AAG	TGC	CTG	GCC	AAT	GGC	TCC	TGG	ACA	GAT	ATG	GAC	ACA	CCC	288
	H18	65-1	wo	SEQ		<del></del>			58			-		<u></u>			

	Val	Arg	Lys	Cys	Leu	Ala	Asn	Gly	Ser	Trp	Thr	Asp	Met	Asp	Thr	Pro	
					85					90					95		
	AGC	CGC	TGT	GTC	CGA	ATC	TGT	TCC	AAG	TCA	TAT	TTG	GCC	CTG	GAA	ААТ	336
5	-			_		Ile							_				
				100			-		105		-			110			
	GGG	AAG	GTC	TTC	CTG	ACG	GGT	GGG	GAC	CTC	CCC	GCT	CTG	GAT	GGA	GCC	384
	Gly	Lys	Val	Phe	Leu	Thr	Gly	Gly	Asp	Leu	Pro	Ala	Leu	Asp	Gly	Ala	
10			115					120					125				
						TGT											432
	Arg		Asp	Phe	Arg	Суѕ	_	Pro	Asp	Phe	His		Val	Gly	Ser	Ser	
		130					135					140					
15	CCG	አርጥ	አጥሮ	un Caur	እርጥ	CAG	ccc	CAG	TCC	) CC	እርጥ	ccc	7 7 C	ccc	CAC	TCC	480
						Gln											400
	145	001		Cyo		150	<b>-</b>	<b></b>			155	110	2,5	110	1113	160	
	-																
20	CAG	GTG	AGC	CGA	ACG	CCG	CAC	TCA	GAG	CGG	CGA	GCG	GTG	TAC	ATC	GGG	528
	Gln	Val	Ser	Arg	Thr	Pro	His	Ser	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	
					165					170					175		
	GCG	CTG	TTT	CCC	ATG	AGC	GGG	GGC	TGG	CCG	GGG	GGC	CAG	GCC	TGC	CAG	576
25	Ala	Leu	Phe		Met	Ser	Gly	Gly	_	Pro	Gly	Gly	Gln		Суз	Gln	
				180					185					190			
	ccc	ccc	cmc	CAC	N MC	GCG	cmc	CAC	CAC	CITIC	3 3 M	100	000	300	CAC	<b>3.00</b> C	624
						Ala											024
30	110	u	195	014	1.00	,,,,,	<b>500</b>	200	p	•	11,511	Jei	205	y	nop	110	
	CTG	CCG	GAC	TAC	GAG	CTC	AAG	CTC	ATC	CAC	CAC	GAC	AGC	AAG	TGT	GAC	672
	Leu	Pro	Asp	Tyr	Glu	Leu	Lys	Leu	Ile	His	His	Asp	Ser	Lys	Cys	Asp	
		210					215					220					
35																	
	CCA	GGC	CAA	GCT	ACC	AAG	TAC	CTG	TAT	GAA	CTG	CTC	TAC	AAC	GAC	CCC	720
	Pro	Gly	Gln	Ala	Thr	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro	
	225					230					235					240	
																	760
40						ATG											768 ر
	Ile	Lys	ile	ile		Met	Pro	GIÀ	Cys		ser	val	Ser		Leu 255	vaı	
					245					250					233		

								•	•				•				
	GCT	GAG	GCT	GCC	AGG	ATG	TGG	AAC	CTC	ATT	GTG	CTC	TĆC	TAT	GGT	TCC	816
	Ala	Glu	Ala	Ala	Arg	Met	Trp	Asn	Leu	Ile	Val	Leu	Ser	Tyr	Gly	Ser	
				260					265					270			
5	AGC	TCA	CCA	GCT	CTG	TCC	AAC	CGG	CAG	CGC	TTT	CCT	ACC	TTC	TTC	CGA	864
	Ser	Ser	Pro	Ala	Leu	Ser	Asn	Arg	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	
			275					280					285				
	ACT	CAT	ccc	TCG	GCC	ACG	CTC	CAC	AAC	ССТ	ACG	CGA	GTG	AAG	CTC	TTT	912
10	Thr	His	Pro	Ser	Ala	Thr	Leu	His	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe	
		290					295					300					
	GAG	AAG	TGG	GGC	TGG	AGG	AAG	ATT	GCC	ACC	ATC	CAG	CAG	ACC	ACC	GAG	960
	Glu	Lys	Trp	Gly	Trp	Arg	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	
15	305					310					315					320	
	GTG	TTC	ACA	TCG	ACT	CTG	GAC	GAC	CTA	GAG	GAA	CGA	GTG	AAG	GAG	GCT	1008
	Val	Phe	Thr	Ser	Thr	Leu	Asp	Asp	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala	
					325					330					335		
20																	
	GGG	ATT	GAG	ATT	ACT	TTC	CGC	CAG	AGC	TTC	TTC	TCA	GAT	CCT	GCC	GTG	1056
	Gly	Ile	Glu	Ile	Thr	Phe	Arg	Gln	Ser	Phe	Phe	Ser	Asp	Pro	Ala	Val	
				340					345					350			
25	CCT	GTC	AAG	AAC	CTC	AAG	CGC	CAG	GAT	GCC	CGA	ATC	ATC	GTG	GGA	CTT	1104
	Pro	Val	Lys	Asn	Leu	Lys	Arg	Gln	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	
			355					360					365				
															AAG		1152
30	Phe	Tyr	Glu	Thr	Glu	Ala	Arg	Lys	Val	Phe	Cys	Glu	Val	Tyr	Lys	Glu	
		370					375					380					
															TAT		1200
	_	Leu	Phe	Gly	Lys	_	Tyr	Val	Trp	Phe		Ile	Gly	Trp	Tyr		
35	385					390					395					400	
	~		me-	m= -			m> -	03.5		ma-	x		ma-			<b>~</b>	1040
															GTG		1248
	Asp	Asn	'l'rp	Phe		Thr	туr	Asp	Pro		тте	ASN	cys	TNY	Val	Asp	
					405					410					415		
40	<b></b> -			as -			<b>63</b> -		<b>~</b>	3 m~	200	) am	<b>~</b>	y mm	cm-	አመሮ	υ 1296
															GTC		1296
	Glu	Met	Thr		Ala	Val	Glu	GIA		TTE	Thr	ınr	GIU		Val	Met	
				420					425					430			

	7710	<i>(</i>	WO	CEC													
	nλa	neu	FIIG	116	261	Val	261	Val	neu	Jer	Jei	Den	GTÅ	116	ACT	Ter	
			TTC Phe														1824
			-							mc				·			1004
40				580					585					590			υ
	Pro	Ala	Asp	Gln	Thr	Leu	Val	Ile	Lys	Thr	Phe	Arg	Phe	Met	Ser	Gln	
	CCG	GCC	GAC	CAG	ACC	CTG	GTC	ATC	AAG	ACA	TTT	CGC	TTC	ATG	TCA	CAG	1776
-		- 5-	_		565		• -	<b>-</b>	- &	570			-3		575		
35			Leu														
	GAT	GAC	CTT	TCC	TGG	тст	AAA	ACG	GAC	AAA	TGG	АТТ	GGA	GGG	GCC	CCC	1728
	545					550					555					560	
			Gly	Gly	Ser		Lys	Lys	Ile	Gly		Tyr	Asp	Ser	Thr		
30			GGT										-	-			1680
		530					535					540					
	_		Asp	_		_											
~	GTG	TTT	GAT	GCC	AGC	GGC	TCA	CGG	ATG	GCC	TGG	АСТ	CTG	ATT	GAG	CAG .	1632
25			212					J2U					525				
	Tyr	Arg	Ala 515	Met	Asn	Ser	Ser	Ser 520	Phe	Glu	Gly	Val		Gly	His	Val	
			GCA														1584
20				500					505					510			
			Glu														
	CGC	CTG	GAA	GAC	TTC	AAC	TAC	AAC	AAC	CAG	ACG	ATC	ACA	GAC	CAA	ATC	1536
					485					490					495		
15	Ala	Leu	Ala	Leu		Lys	Thr	Ser	Gly	_	Ser	Gly	Arg	Ser	_	Val	
			GCC														1488
	465					470					475					480	
			Phe														
10	GGC	GGC	TTC	CAG	GAG	GCA	CCG	CTG	GCC	TAT	GAT	GCC	ATC	TGG	GCC	TTG	1440
		450					#33					460					
	Phe	Val 450	Glu	гÀ2	Leu	Thr	Lys 455	Arg	Leu	rys	Arg		Pro	Glu	Glu	Thr	
			GAG														1392
5																	
			435					440					445				
	Leu	Asn	Pro	Ala	Asn	Thr	Arg	Ser	Ile	Ser	Asn	Met	Thr	Ser	Gln	Glu	
	CTG	AAC	CCA	GCC	AAC	ACC	CGC	AGC	ATC	TCC	AAC	ATG	ACA	TCC	CAG	GAG	1344
													•				· <del></del>

	H18	865-1	wo	SEQ				(	52			_	<del></del>				
	ATG	AAC	ACC	TGG	CTT	GGC	ATT	TTC	TAT	GGT	TAC	AAG	GGG	CTG	CTG	CTG	2352
			755					760					765				
40											His						
	ATT	GAT	GTG	TCC	ATC	CTG	ccc	CAG	CTG	GAG	CAC	TGC	AGC	TCC	AAG	AAA	2304
			J	740					745	•				- 750		_	
<b>,,</b>											Glu						2200
35	באנים	CAC	CGG	ACC		GAG	ልሮጥ	փփո	GCC	AAG	GAG	GAA	CCA	AAG	GAA	GAT	2256
	Val	Gly	Met	Asp	Val 725	Leu	Thr	Leu	Ala	Tle 730	Trp	Gln	Met	Val	Asp 735	Pro	
			ATG								TGG						2208
30						-											
	705	гуз	rnr	ьeu	GIU	710	тф	гЛя	neu	TAL	715	Inr	vaı	GTĀ	neu	720	
											ACC Thr						2160
4.0																	
25	Trp	Trp 690	Val	His	Thr	Val	Phe 695	Thr	Lys	Lys	Glu	Glu 700	Lys	Lys	Glu	Trp	
											GAG						2112
			3,3					-50									
20	Leu	Gly	Leu 675	GTA	rne	ser	ren	680 GIA	ıyr	СТĀ	Ser	met	Phe 685	Tnr	ьys	TTE	
								_		_	TCC			_		• _	2064
	Ile	Gly	Arg	Ser 660	Gln	Phe	Pro	Phe	Val 665	Суѕ	Gln	Ala	Arg	Leu 670	Trp	Leu	
15											CAG						2016
					043					020					033		
	Ser	Leu	Ala	Leu	Ala 645	Ala	Val	Phe	Pro	Leu 650	Gly	Leu	Asp	Gly	Tyr 655	His	,
		-									GGG			•			1968
10																	
	625	GIN	ASN	ser	GIII	630	ASII	neu	ASII	ASII	Leu 635	1111	ATA	vai	GIŞ	640	
											CTG				_		1920
5		610					615					620					
	Ala		Val	Cys	Leu	Ser		Asn	Ile	Tyr	Asn		His	Val	Arg	Tyr	
	GCT	GTG	GTC	TGT	CTG	TCC	TTT	AAC	ATC	TAC	AAC	TCT	CAT	GTC	CGT	TAC	1872
			595					600				٠	605				
								•	-				•				

SUBSTITUTE SHEET (RULE 26)

									-								
	Met	Asn	Thr	Trp	Leu	Gly	Ile	Phe	Tyr	Gly	Tyr	Lys	Gly	Leu	Leu	Lėū	
		770					775					780					
	CMC	CULY	ccc	א תיכי	danda.	Cum	CCT	ጥልጥ	CAC	ACC	AAG	) AGC	GTG.	ጥርጥ	ACT	CAC	2400
5													_	_	Thr	_	2400
J	785	Dea	O+y	110	1110	790	7124	-3-	014		795	-				800	
	AAG	ATC	AAT	GAC	CAC	CGG	GCT	GTG	GGC	ATG	GCC	ATG	TAC	AAC	GTG	GCG	2448
	Lys	Ile	Asn	Asp	His	Arg	Ala	Val	Gly	Met	Ala	Met	Tyr	Asn	Val	Ala	
10					805					810					815		
																	2426
															AGC	_	2496
	Val	Leu	суѕ	820	TIE	Thr	Ala	Pro	825	Thr	met	ire	Leu	830	Ser	GIN	
15				020					023					030			
••	CAG	GAT	GCA	GCT	TTC	GCC	TTT	GCA	GCT	CTT	GCC	ATA	GTG	TTC	TCC	TCC	2544
	Gln	Asp	Ala	Ala	Phe	Ala	Phe	Ala	Ala	Leu	Ala	Ile	Val	Phe	Ser	Ser	
			835					840					845				
										ر							
20															TTG		2592
	Tyr		Thr	Leu	Val	Val		Phe	Val	Pro	Lys		Arg	Arg	Leu	Ile	
		850					855					860					
	ACC	CGG	GGT	GAG	TGG	CAG	TCG	GAG	GCG	CAG	GAT	ACC	ATG	AAA	ACG	GGG	2640
25	Thr	Arg	Gly	Glu	Trp	Gln	Ser	Glu	Ala	Gln	Asp	Thr	Met	Lys	Thr	Gly .	
	865					870					875					880	
	TCG	TCG	ACC	AAC	AAC	AAT	GAG	GAA	GAG	AAG	TCC	CGA	CTG	TTG	GAG	AAG	2688
	Ser	Ser	Thr	Asn		Asn	Glu	Glu	Glu	_	Ser	Arg	Leu	Leu	Glu	Lys	
30					885					890					895		
	GAG	AAC	CGG	GAG	СТС	GAG	AAG	ATC	АТТ	GCT	GAG	AAA	GAG	GAG	CGA	GTG	2736
															Arg		
			_	900			_		905					910			
35																	
	TCC	GAG	CTG	CGC	CAT	CAG	CTT	CGT	TCT	CGG	CAG	CAG	CTG	CGC	CCT	CGG	2784
	Ser	Glu	Leu	Arg	His	Gln	Leu	Arg	Ser	Arg	Gln	Gln	Leu	Arg	Pro	Arg	
			915					920					925				
								<b>~</b> -~	000	ma:	000	000	080	000	200	CON	2022
40															AGG	GGA	2832 <sub>v</sub>
	Arg	930		rro	THE	PIO	935	vab	FIO	PET	GIY	940	nen	FIO	,	OLY.	
		930					ر د ر				•						

CCC CAT GAG CCC CCT GAC CGG CTC AGC TGT GAC GGG AGC CGG GTT CAC

Pro His Glu Pro Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His

945

950

960

TTG CTG TAC AAG TGA

2895

Leu Leu Tyr Lys \*

965

- 10 (2) INFORMATION FOR SEQ ID NO: 53:
  - (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 964 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:

20

35

Met Leu Leu Leu Leu Pro Leu Ala Leu Ala Pro Leu Phe Leu Arg

1 5 10 15

Pro Pro Gly Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly
25 20 25 30

Cys Gln Ile Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly
35 40 45

30 Leu Thr Arg Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr 50 55 60

Glu Ile Glu Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys
65 70 75 80

Val Arg Lys Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro

Ser Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Ala Leu Glu Asn 40 100 105 110

Gly Lys Val Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala 115 120 125

	Arg	Val	Asp	Phe	Arg	Cys	Asp	Pro	Asp	Phe	His	Leu	Val	Gly	Ser	Ser
		130					135					140				
5	Arg	Ser	Ile	Cys	Ser	Gln	Gly	Gln	Trp	Ser	Thr	Pro	Lys	Pro	His	Cys
	145					150					155					160
	Gln	Val	Ser	Arg	Thr	Pro	His	Ser	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly
					165					170					175	
10																
	Ala	Leu	Phe	Pro	Met	Ser	Gly	Gly	Trp	Pro	Gly	Gly	Gln	Ala	Cys	Gln
				180					185					190		
	Pro	Ala	Val	Glu	Met	Ala	Leu	Glu	Asp	Val	Asn	Ser	Arg	Arg	Asp	Ile
15			195					200					205		_	
	Leu	Pro	Asp	Tyr	Glu	Leu	Lys	Leu	Ile	His	His	Asp	Ser	Lys	Cys	Asp
		210					215					220				
20	Pro	Gly	Gln	Ala	Thr	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro
	225					230					235					240
	Ile	Lys	Ile	Ile	Leu	Met	Pro	Gly	Cys	Ser	Ser	Val	Ser	Thr	Leu	Val
					245					250					255	
25																
	Ala	Glu	Ala	Ala	Arg	Met	Trp	Asn	Leu	Ile	Val	Leu	Ser	Tyr	Gly	Ser
				260					265					270		
	Ser	Ser	Pro	Ala	Leu	Ser	Asn	Arg	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg
30			275					280					285			
	Thr	His	Pro	Ser	Ala	Thr	Leu	His	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe
		290					295					300				
35	Glu	Lys	Trp	Gly	Trp	Arg	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu
	305					310					315					320
	Val	Phe	Thr	Ser	Thr	Leu	Asp	Asp	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala
					325					330					335	
40																
	Gly	Ile	Glu	Ile	Thr	Phe	Arg	Gln	Ser	Phe	Phe	Ser	Asp	Pro	Ala	Val
				340					345					350	•	

Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Thr Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu 25 Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Ser Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ala Pro Pro Ala Asp Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Met Ser Gln

Lys Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Ser Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys Leu Tyr Thr Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Met Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Lys Lys 

Met Asn Thr Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu 770 775 780

Leu Leu Gly Ile Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu
785 790 795 800

Lys Ile Asn Asp His Arg Ala Val Gly Met Ala Met Tyr Asn Val Ala 805 810 815

Val Leu Cys Leu Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln 820 Gln Asp Ala Ala Phe Ala Phe Ala Leu Ala Ile Val Phe Ser Ser 840 Tyr Ile Thr Leu Val Val Leu Phe Val Pro Lys Met Arg Arg Leu Ile 850 855 860 10 Thr Arg Gly Glu Trp Gln Ser Glu Ala Gln Asp Thr Met Lys Thr Gly 870 875 Ser Ser Thr Asn Asn Asn Glu Glu Glu Lys Ser Arg Leu Leu Glu Lys 885 890 15 Glu Asn Arg Glu Leu Glu Lys Ile Ile Ala Glu Lys Glu Glu Arg Val 900 905 Ser Glu Leu Arg His Gln Leu Arg Ser Arg Gln Gln Leu Arg Pro Arg 915 920 925 Arg His Pro Pro Thr Pro Pro Asp Pro Ser Gly Gly Leu Pro Arg Gly 930 935 940 25 Pro His Glu Pro Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His 945 950 955 960 Leu Leu Tyr Lys 30 (2) INFORMATION FOR SEQ ID NO: 54: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1737 base pairs 35 (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: cDNA to mRNA 40

H1865-1 WO SEQ

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens 5 (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1..1737 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 54: 10 ATG TTG CTG CTG CTA CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC 48 Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly 5 10 15 15 GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC 96 Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile 20 25 ATA CAC CCG CCC TGG GAA GGG GGC ATC AGG TAC CGG GGC CTG ACT CGG 144 Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg 35 GAC CAG GTG AAG GCT ATC AAC TTC CTG CCA GTG GAC TAT GAG ATT GAG 192 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 TAT GTG TGC CGG GGG GAG CGC GAG GTG GTG GGG CCC AAG GTC CGC AAG 240 Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys 70 65 75 30 TGC CTG GCC AAC GGC TCC TGG ACA GAT ATG GAC ACA CCC AGC CGC TGT 288 Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys 35 GTC CGA ATC TGC TCC AAG TCT TAT TTG ACC CTG GAA AAT GGG AAG GTT 336 Val Arg Ile Cys Ser Lys Ser Tyr Leu Thr Leu Glu Asn Gly Lys Val 100 TTC CTG ACG GGT GGG GAC CTC CCA GCT CTG GAC GGA GCC CGG GTG GAT 384, Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala Arg Val Asp 115 120 125

	7710	<i>CE</i> 1	1110	050					70								
		270					2,7,3					200					
	Ser	290	IIII	neu	ura	Vali	295	TITE	nr 9	Val	בעעם	300	I IIG	<b></b>	Lys	F	
															Lys		
40	ጥሮል	GCC	ACA	Curc	ראכ	220	ССт	ልሮሮ	כפר	GTG	ΔΔΔ	כתר	Тфф	GAA	AAG	TGG	912
40			213					200					<b>403</b>			•	
	WIG	Leu		usii	Arg	GIII	vrā	280	FIU	1111	FILE	LIIG	285	1111	1173		
															His		304
	ccc	CTC	ጥርን	אאר	ccc	CNG	CCm	መመር	ccc	א רייי	danc.	ιτιπ <b>ι</b>	CGA	) )	CAC	CCA	864
35				260					265					270			
25	ATS	AIG	met	_	ASN	ьeu	тте	vai		ser	ıyr	стλ	ser		Ser	PEO	
															TCA		816
	~~	100	<b>.</b>	mc~		OE C	3 mm	ome.		maa	m - m	000	maa		m~3	CCA	03.6
					245					250					255		
30	Ile	Leu	Met	Pro		Cys	Ser	Ser	Val		Thr	Leu	Val	Ala	Glu	Ala	
															GAG	_	768
																	_
	225					230					235					240	
		Thr	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro	Ile	Lys	Ile	
25	GCC	ACC	AAG	TAC	CTA	TAT	GAG	CTG	CTC	TAC	AAC	GAC	CCT	ATC	AAG	ATC	720
		210					215					220					
	Tyr	Glu	Leu	Lys	Leu	Ile	His	His	Asp	Ser	Lys	Cys	Asp	Pro	Gly	Gln	
	TAT	GAG	CTC	AAG	CTC	ATC	CAC	CAC	GAC	AGC	AAG	TGT	GAT	CCA	GGC	CAA	672
20																	
			195					200					205				
	Glu	Met	Ala	Leu	Glu	Asp	Val	Asn	Ser	Arg	Arg	Asp	Ile	Leu	Pro	Asp	
	GAG	ATG	GCG	CTG	GAG	GAC	GTG	AAT	AGC	CGC	AGG	GAC	ATC	CTG	CCG	GAC	624
15			. =	180	- 4	. •		-4	185	. =				190		<del></del>	
				_	_			_	_	_	_				Ala		2.0
	CCC	ATG	AGC	GGG	GGC	TGG	CCA	GGG	GGC	CAG	GCC	ፐርር	CAG	CCC	GCG	GTG	576
					103					110					175		
10	Arg	rnr	PTO	nis	165	GIU	Arg	Arg	AIA	Val 170	ıyr	тте	GIĀ	ALA	Leu 175	hue	
10															CTG		528
		100	CCN	a.c	ma v	C 2 2 2	000	000	CC.	CMC	ma c	<b>3</b> mc	600	201	ama	mmm	520
	145					150					155					160	
	_	Ser	Gln	Gly	Gln		Ser	Thr	Pro	Lys		His	Суѕ	Gln	Val		
5												_			GTG		480
		130					135					140					
	Phe	Arg	Cys	Asp	Pro	Asp	Phe	His	Leu	Val	Gly	Ser	Ser	Arg	Ser	Ile	
	TTC	CGG	TGT	GAC	ccc	GAC	TTC	CAT	CTG	GTG	GGC	AGĊ	TCC	CGĠ	AGC	ATC	432
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	GIN	GIU	ATA	rro	Leu	ATA	ıyr	ASP	ATA	тте	Trp	AIA	ьeи	AIA	ren	AT9		
					CTG					*							1440	)
40		450					455					460						U
	Lys	Leu	Thr	Lys	Arg	Leu	Lys	Arg	His	Pro	Glu	Glu	Thr	Gly	Gly	Phe	•	
	AAA	CTA	ACC	AAG	CGA	CTG	AAA	AGA	CAC	CCT	GAG	GAG	ACA	GGA	GGC	TTC	139	2
33	VIO	NOII	435	n. y	261	115	Jer	440	1.1CC	1111	Jer	GIII	445	FIIG	val	GIU		
35					Ser												1344	•
	ccc	חתממ	እሮሮ	ccc	AGC	יוויניו ע	TICC	220	አጥር	מסמ	TCC	CVC	CAA	mmm	CITIC	CAC	124	4
				420					425					430				
	Glu	Ala	Val		Gly	His	Ile	Thr		Glu	Ile	Val	Met		Asn	Pro		
30	GAG	GCG	GTG	GAG	GGC	CAC	ATC	ACA	ACT	GAG	ATT	GTC	ATG	CTG	AAT	CCT	129	5
				-1-	405	•				410				J14	415	****		
	_		<u>.</u>		GAC Asp			_									124	3
25	ጥጦጣ	አአሮ	איייע	መአብ	GAC	CCm	ηn,c.m	አ መ ଦ	ח א א	THE C	202	Cmc	Cym	CAC	አሙግ	a.cm	. 194	٥
	385					390					395					400		
	Gly	Lys	Lys	Tyr	Val	Trp	Phe	Leu	Ile	Gly	Trp	Tyr	Ala	Asp	Asn	Trp		
	GGG	AAG	AAG	TAC	GTC	TGG	TTC	CTC	АТТ	GGG	TGG	TAT	GCT	GAC	AAT	TGG	120	C
20		370		9	~, 3		375	<b>-</b> 13	u	- W.L	-7.	380	U.L.	ni 9	reu	1116		
					Lys						_						. 115	۷
	N C/FF	C2.2	ccc	ccc	AAA	Cmm	птт	mom.	CAC	CEC	ma c	220	C 2 C	~~~	Omc.	(D/DP)	335	,
			355					360					365					
15	Asn	Leu	_	Arg	Gln	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	Phe	Tyr	Glu		
	AAC	CTG	AAG	CGC	CAG	GAT	GCC	CGA	ATC	ATC	GTG	GGA	CTT	TTC	TAT	GAG	110	4
	**6	****		340		Jer			345	ענה	110	ard	Val	350	AGT	- Py S		
10					Gln												105	3
10	ልጣጣ	ልርመ	ሙጥ	CGC	CAG	ልርመ	ጥጥ	ጥጥረ	тсх	Gam	CCA	CCT	CITIC	ccc	GMC	227	105	5
					325					330					335			
	Ser	Thr	Leu	qaA	Asp	Leu	Glu	Glu	Arg		Lys	Glu	Ala	Gly		Glu		
	TCG	ACT	CTG	GAC	GAC	CTG	GAG	GAA	CGA	GTG	AAG	GAG	GCT	GGA	ATT	GAG	100	8
5																		
	305					310					315					320		
					Ile												- •	
	GGC	TGG	AAG	AAG	АТТ	GCT	ACC	ATC	CAG	CAG	ACC	ACT	GAG	GTC	TTC	ACT	96	0
												•	•	•			_	

	wo	99/2	1890						-						PCT/SE98/0	1947
	465					470				475					480	-
<b>5</b> .										TCT Ser						1488
										GAC Asp						1536
10										GGC						1584
15										ATC Ile						1632
20						_	_			AGC Ser 555						1680
25										ATA Ile						1728
		ACC Thr	TGA *													1737
30	(2)	INFO	ORMA	rion	FOR	SEQ	ID I	 55:								
35			(1	A) L:	engti Ype :	H: 5' ami										
40							pro:	SEQ :	ID NO	D: 5!	5:					,
	Met									Phe		Arg	Pro	Pro	Gly	-

72

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· 1 Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys 15 Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Thr Leu Glu Asn Gly Lys Val Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala Arg Val Asp Phe Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser Arg Ser Ile Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys Gln Val Asn Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile 

	Ile	Leu	Met	Pro	Gly 245	Cys	Ser	Ser	Val	Ser 250	Thr	Leu	Val	Ala	Glu 255	Ala
5	Ala	Arg	Met	Trp 260	Asn	Leu	Ile	Val	Leu 265	Ser	Tyr	Gly	Ser	Ser 270	Ser	Pro
10	Ala	Leu	Ser 275	Asn	Arg	Gln	Arg	Phe 280	Pro	Thr	Phe	Phe	Arg 285	Thr	His	Pro
	Ser	Ala 290	Thr	Leu	His	Asn	Pro 295	Thr	Arg	Val	Lys	Leu 300	Phe	Glu	Lys	Trp
15	Gly 305	Trp	Lys	Lys	Ile	Ala 310	Thr	Ile	Gln	Gln	Thr 315	Thr	Glu	Val	Phe	Thr 320
	Ser	Thr	Leu	Asp	Asp 325	Leu	Glu	Glu	Arg	Val 330	Lys	Glu	Ala	Gly	Ile 335	Glu
20	Ile	Thr	Phe	Arg 340	Gln	Ser	Phe	Phe	Ser 345	Asp	Pro	Ala	Val	Pro 350	Val	Lys
25	Asn	Leu	Lys 355	Arg	Gln	Asp	Ala	Arg 360	Ile	Ile	Val	Gly	Leu 365	Phe	Туг	Glu
	Thr	Glu 370	Ala	Arg	Lys	Val	Phe 375	Cys	Glu	Val	Туг	Lys 380	Glu	Arg	Leu	Phe
30	Gly 385	Lys	Lys	Tyr	Val	Trp 390	Phe	Leu	Ile	Gly	Trp 395	Tyr	Ala	Asp	Asn	Trp 400
	Phe	Lys	Ile	туг	Asp 405	Pro	Ser			-			Asp		Met 415	Thr
35	Glu	Ala	Val	Glu 420	Gly	His	Ile	Thr	Thr 425	Glu	Ile	Val	Met	Leu 430	Asn	Pro
40	Ala	Asn	Thr 435	Arg	Ser	Ile	Ser	Asn 440	Met	Thr	Ser	Gln	Glu 445	Phe	Val	Glu
70	Lys	Leu 450	Thr	Lys	Arg	Leu	Lys 455	Arg	His	Pro	Glu	Glu 460	Thr	Gly	Gly	Phe

Gin Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala 470 475 Leu Asn Lys Thr Ser Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu 485 490 Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala 500 505 510 Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp 515 520 525 Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly 535 540 530 15 Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu 545 550 555 560 Ser Trp Ser Lys Thr Asp Lys Trp Ile Val Ile Ser Arg Thr His Ser 570 565 575 20 Pro Thr 25 (2) INFORMATION FOR SEQ ID NO: 56: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1386 base pairs (B) TYPE: nucleic acid 30 (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: cDNA to mRNA 35 (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens

(ix) FEATURE:

(A) NAME/KEY: CDS

(B) LOCATION:1..1386

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 56:

		(xi)	SEC	QUENC	CE DE	ESCRI	PTIC	ON: S	SEQ I	D NO	): 56	5:					
5																	
	ATG	GGG	CCC	GGG	GCC	CCT	TTT	GCC	CGG	GTG	GGG	TGG	CCA	CTG	CCG	CTT	48
	Met	Gly	Pro	Gly	Ala	Pro	Phe	Ala	Arg	Val	Gly	Trp	Pro	Leu	Pro	Leu	
	1				5					10					15		
10	CTG	GTT	GTG	ATG	GCG	GCA	GGG	GTG	GCT	CCG	GTG	TGG	GCC	TCC	CAC	TCC	96
	Leu	Val	Val	Met	Ala	Ala	Gly	Val	Ala	Pro	Val	Trp	Ala	Ser	His	Ser	•
				20					25					30			•
	CCC	CAT	CTC	CCG	CGG	CCT	CAC	TCG	CGG	GTC	CCC	CCG	CAC	CCC	TCC	TCA	144
15	Pro	His	Leu	Pro	Arg	Pro	His	Ser	Arg	Val	Pro	Pro	His	Pro	Ser	Ser	
			35					40					45				
	GAA	CGG	CGC	GCA	GTG	TAC	ATC	GGG	GCA	CTG	TTT	CCC	ATG	AGC	GGG	GGC	192
	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	Ala	Leu	Phe	Pro	Met	Ser	Gly	Gly	
20		50					55					60					
				GGC													240
	Trp	Pro	Gly	Gly	Gln	Ala	Суѕ	Gln	Pro	Ala	Val	Glu	Met	Ala	Leu	Glu	
	65					70					75					80	
25																	
				AGC									_				288
	Asp	Val	Asn	Ser	_	Arg	Asp	Ile	Leu		Asp	Tyr	Glu	Leu		Leu	
					85					90					95		
																	226
30				GAC													336
	Ile	His	Hls	Asp	Ser	ьуs	Cys	Asp		GIĀ	GIN	Ala	THE		туг	Leu	
				100					105					110			
	<b></b>	G1.G	ama	CTC	mag	220	CAC	CCM	3.000	220	» mc	» mc	C/II/III	אתכ	CCM	ccc	384
25																	204
35	туг	GIU		Leu	TYL	ASII	Asp	120	116	Dys	TIE	116	125	Hec	FIO	GLY	
			115					120					123				
	mcc.	200	mcm	CMC	mcc.	ACC.	כיזיכ	CTC	CCT	GAG	CCT	CCT	ACC	אתכ	TGG	AAC	432
																Asn	
40	Cys	130	Ser	Val	Ser	1111	135	Vul	nia	014	1114	140	.u.g	1100			•
40		T30										~40					υ
	CITIC	שתו ע	GTVC	Cmm	ייירר	ጥልጥ	GGC	ጥርር	AGC	ጥሮል	CC2	GCC	ርጥር	ጥሮል	AAC	CGG	480
																Arg	-50
	Leu	116	val	neu	361	+ 3 -	OLY	261	DCT		0						

	wo	99/2	1890							-						PCT/S	SE98/01947
	145					150					155		ē	٠		160	_
	CAG	CGT	TTC	ccc	ACT	TTC	TTC	CGA	ACG	CAC	CCA	TCA	GCC	ACA	CTC	CAC	528
	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	His	Pro	Ser	Ala	Thr	Leu	His	
5					165					170					175		
	AAC	CCT	ACC	CGC	GTG	AAA	CTC	TTT	GAA	AAG	TGG	GGC	TGG	AAG	AAG	ATT	576
	Asn	Pro	Thr	_	Val	Lys	Leu	Phe	Glu	Lys	Trp	Gly	Trp	Lys	Lys	Ile	
				180					185					190			
10	ር ር ጥ	ACC	ATC	CAG	CAG	ACC	АСТ	GAG	GTC	ጥጥር	ACT	ጥርር	ACT	CTG	GAC	GAC	624
											Thr						
			195					200					205				
15	CTG	GAG	GAA	CGA	GTG	AAG	GAG	GCT	GGA	ATT	GAG	ATT	ACT	TTC	CGC	CAG	672
	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala	Gly	Ile	Glu	Ile	Thr	Phe	Arg	Gln	
		210					215					220					
	».cm	mmc	mmc	mc s	CAM	CCA	ccm	cmc	ccc	CMC	222	220	cmc	***	ccc	CNC	720
20											AAA Lys						720
20	225	riie	rne	Ser	АЗР	230	AIG	vai	PLO	Val	235	ASII	Leu	гуs	ALG	240	
	223															2.10	
	GAT	GCC	CGA	ATC	ATC	GTG	GGA	CTT	TTC	TAT	GAG	ACT	GAA	GCC	CGG	AAA	768
	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	Phe	Tyr	Glu	Thr	Glu	Ala	Arg	Lys	
25					245					250					255		
	GTT	TTT	TGT	GAG	GTG	TAC	AAG	GAG	CGT	CTC	TTT	GGG	AAG	AAG	TAC	GTC	816
	Val	Phe	Cys		Val	Tyr	Lys	Glu	-	Leu	Phe	Gly	Lys	-	Tyr	Val	
				260					265					270			
30	TGG	ጥጥር	ርጥር	<b>አ</b> ጥጥ	GGG	TYCG	ייע	CCT	GAC	<u>አ</u> ልጥ	TGG	ጥጥር	AAG	ልጥሮ	ጥልሮ	GAC	864
											Trp	_					
			275			-	-	280			-		285		•	•	
35	CCT	TCT	ATC	AAC	TGC	ACA	GTG	GAT	GAG	ATG	ACT	GAG	GCG	GTG	GAG	GGC	912
	Pro		Ile	Asn	Суѕ	Thr		Asp	Glu	Met	Thr		Ala	Val	Glu	Gly	
		290					295			,		300					
	CAC	ልጥር	aca	ልሮጥ	GAG	ልጥጥ	GTYC	ልጥሮ	ריויני	יתמ	ССТ	GCC	ልልጥ	<b>ACC</b>	CGC	AGC	960
40											Pro						1
,,,	305					310					315				- 3	320	1
															•		
	ATT	TCC	AAC	ATG	ACA	TCC	CAG	GAA	TTT	GTG	GAG	AAA	CTA	ACC	AAG	CGA	1008

CTG AAA AGA CAC CCT GAG GAG ACA GGA GGC TTC CAG GAG GCA CCG CTG  Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu  340 345 350  GCC TAT GAT GCC ATC TGG GCC TTG GCA CTG GCC CTG AAC AAG ACA TCT  Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser  360 365 365  GGA GGA GGC GGC CGT TCT GGT GTG CGC CTG GAG GAC TTC AAC TAC AAC  Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn  370 375 380   AAC CAG ACC ATT ACC GAC CAA ATC TAC CGG GCA ATG AAC TCT TCG TCC  Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser  385 390 395 400  TTT GAG GGT GTC TCT GGC CAT GTG GTG TTT GAT GCC AGC GGC TCT CGG  Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg  405 410 415  ATG GCA TGG ACG CTT ATC GAG CAG CAG GTT CAG GGT GGC AGC TAC AAG AAG  25 Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys  420 425 425  ATT GGC TAC TAT GAC AGC ACC AAG GAT GAT CTT TCC TGG TCC AAA ACA  Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr  435 440 445  GAT AAA TGG ATT GTT ATA TCC AGA ACC CAC AGC CCA ACC TGA  Asp Lys Trp Ile Val Ile Ser Arg Thr His Ser Pro Thr *  450 455 455		Ile	Ser	Asn	Met	Thr 325	Ser	Gln	Glu	Phe	Val 330	Glu	Lys	Leu	Thr	Lys 335	Arg		
Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser    10	5				His					Gly					Ala				1056
Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn 370	10			Asp					Leu					Asn					1104
AAC CAG ACC ATT ACC GAC CAA ATC TAC CGG GCA ATG AAC TCT TCG TCC ASN Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser 385			Gly					Gly					Asp						1152
Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg 405	15	Asn					Asp					Ala					Ser		1200
Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys 420	20					Ser					Phe					Ser			1248
Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr  440  GAT AAA TGG ATT GTT ATA TCC AGA ACT CAC AGC CCA ACC TGA  Asp Lys Trp Ile Val Ile Ser Arg Thr His Ser Pro Thr  450  455  460	25				Thr					Leu					Tyr			·	1296
Asp Lys Trp Ile Val Ile Ser Arg Thr His Ser Pro Thr * 450 455 460	30			Tyr					Lys				-	Trp					1344
	35		Lys					Ser					Pro	_	TGA *				1386

(2) INFORMATION FOR SEQ ID NO: 57:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 461 amino acids

(B) TYPE: amino acid(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:

Met Gly Pro Gly Ala Pro Phe Ala Arg Val Gly Trp Pro Leu Pro Leu

10 15

Leu Val Val Met Ala Ala Gly Val Ala Pro Val Trp Ala Ser His Ser
20 25 30

10 Pro His Leu Pro Arg Pro His Ser Arg Val Pro Pro His Pro Ser Ser
35 40 45

Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe Pro Met Ser Gly Gly
50 55 60

Trp Pro Gly Gln Ala Cys Gln Pro Ala Val Glu Met Ala Leu Glu
65 70 75 80

Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu 20 85 90 95

Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala Thr Lys Tyr Leu 100 105 110

25 Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile Leu Met Pro Gly
115 120 125

Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn 130 135 140

Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala Leu Ser Asn Arg 145 150 155 160

Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser Ala Thr Leu His 35 165 170 175

Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly Trp Lys Lys Ile 180 185 190

40 Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr Ser Thr Leu Asp Asp 195 200 205

Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile Thr Phe Arg Gln

15

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Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Ile Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser 305 -Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys

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Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr 

Asp Lys Trp Ile Val Ile Ser Arg Thr His Ser Pro Thr 450 455 5 (2) INFORMATION FOR SEQ ID NO: 58: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1746 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: cDNA to mRNA (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Canis familiaris (ix) FEATURE: (A) NAME/KEY: CDS 25 (B) LOCATION:1..1746 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 58: 48 Met Leu Leu Leu Leu Pro Leu Ala Leu Ala Pro Leu Phe Leu Arg 5 CCC CCG GGC GCG GGG GCA CAG ACC CCC AAC GCC ACC TCG GAA GGT 96 Pro Pro Gly Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly 35 20 30

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40

35

TGC CAG ATC ATA CAC CCG CCT TGG GAA GGG GGT ATC AGG TAC AGG GGC

Cys Gln Ile Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly

CTG ACT CGT GAC CAG GTG AAG GCT ATC AAC TTC CTG CCG GTG GAC TAT

Leu Thr Arg Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr

144

GAG ATT GAG TAT GTG TGC CGG GGA GAG CGA GAG GTG GTG GGG CCC AAG  Glu Ile Glu Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys 65 70 75 80  70 80  70 80																	- •	
65 70 70 75 80  STO CGA AAG TGC CTG GCC AAT GGC TCC TGG ACA GAT ATG GAC ACA CCC VAIL ATG Lys Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro 95  10 AGC CGC TGT GTC CGA ATC TGT TCC AAG TCA TAT TTG GCC CTG GAA AAT 336  Ser Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Ala Leu Glu Asn 100 105 110  GGG AAG GTC TTC CTG ACG GGT GGG GAC CTC CCC GCT CTG GAA AAT 115 GGC CTG GAA AAT 115 115 120 120 125  CGG GTG GAT TTC CGG TGT GAC CCT GAC TTC CAT CTT GTG GGC AGC TCC AGC AGC AGC AGC AGC AGC AGC AGC AGC A		GAG	АТТ	GAG	TAT	GTG	TGC	CGG	GGA	GAG	CGA	GAG	GTG	GTG	GGG	ccc	AAG	240
GTC CGA AAG TGC CTG GCC AAT GGC TCC TGG ACA GAT ATG GAC ACA CCC  Val Arg Lys Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro 85 90 95  10 AGC CGC TGT GTC CGA ATC TGT TCC AAG TCA TAT TTG GCC CTG GAA AAT Ser Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Ala Leu Glu Asn 100 105 110  GGG AAG GTC TTC CTG ACG GGT GGG GAC CTC CCC GCT CTG GAT GGA GCC 184 115 120 125  CGG GTG GAT TTC CGG TGT GAC CCT GAC TTC CAT CTT GTG GGC AGC TCC Arg Val Asp Phe Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser 130 135 140  CGG AGT ATC TGT AGT CAG GGC CAG TGG AGC ACT CCC AAG CCC CAC TGC Arg Ser Ile Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys 145 150 155 160  25  CAG GTG AGC CGA ACG CCC CAC TCA GAG CGC GGG GGC GTG TAC ATC GGG Gln Val Ser Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly 165 170 175  30 GCG CTG TTT CCC ATG AGC GGG GGC TGG CCG GGG GGC CAG GCC TGC CAG Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180 180 185 195  CCC GCC GTG GAG ATG GCG CTG GAG GCC TGG AAT AGC CCC AAG GAC ATC CCC GCC GTG GAG ATG GCG CTG GAG GCC TGG AAT AGC CCC AAG GAC AGC Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180 185 195  CCC GCC GTG GAG ATG GCG CTG GAG AGC GTG AAT AGC CCC AGG GAC ATC CCC GCC GTG GAG ATG GCG CTG GAG AGC GTG AAT AGC CCC AGG GAC ATC 15 Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile 195 200 200 205  CTG CCG GAC TAC GAG CTC AAG CTC AAC CAC CAC GAC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210 CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720		Glu	Ile	Glu	Tyr	Val	Cys	Arg	Gly	Glu	Arg	Glu	Val	Val	Gly	Pro	Lys	
STC CGA AAG TGC CTG GCC AAT GGC TCC TGG ACA GAT ATG GAC ACA CCC   288		65					70					75					80	
Val Arg Lys Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro	5																	
10   AGC   CGC   TGT   GTC   CGA   ATC   TGT   TCC   AAG   TCA   TAT   TTG   GCC   CTG   GAA   AAT   336   Ser   Arg   Cys   Val   Arg   Ile   Cys   Ser   Lys   Ser   Tyr   Leu   Ala   Leu   Glu   Asn   100   105   110		GTC	CGA	AAG	TGC	CTG	GCC	AAT	GGC	ŢCC	TGG	ACA	GAT	ATG	GAC	ACA	CCC	288
10 AGC CGC TGT GTC CGA ATC TGT TCC AAG TCA TAT TTG GCC CTG GAA AAT 3366 SER Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Ala Leu Glu Asn 100 105 105 110 110 386    GGG AAG GTC TTC CTG ACG GGT GGG GAC CTC CCC GCT CTG GAT GGA GCC 384    15 Gly Lys Val Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala 115 125    CGG GTG GAT TTC CGG TGT GAC CCT GAC TTC CAT CTT GTG GGC AGC TCC A32   Arg Val Asp Phe Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser 130 130		Val	Arg	Lys	Cys	Leu	Ala	Asn	Gly	Ser	Trp	Thr	Asp	Met	Asp	Thr	Pro	
Ser Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Ala Leu Glu Asn   110   100						85					90					95		
Ser Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Ala Leu Glu Asn   110   100																		
GGG   AAG   GTC   TTC   CTG   ACG   GGT   GGG   GAC   CTC   CCC   GCT   CTG   GAT   GGA   GCC   GCT   GIY   Lys   Val   Phe   Leu   Thr   Gly   Gly   Asp   Leu   Pro   Ala   Leu   Asp   Gly   Ala   115   120   125	10	AGC	CGC	TGT	GTC	CGA	ATC	TGT	TCC	AAG	TCA	TAT	TTG	GCC	CTG	GAA	ААТ	336
GGG AAG GTC TTC CTG ACG GGT GGG GAC CTC CCC GCT CTG GAT GGA GCC  15 Gly Lys Val Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala 115 120 125 25 25 25 25 25 25 25 25 25 25 25 25 2		Ser	Arg	Cys	Val	Arg	Ile	Cys	Ser	Lys	Ser	Tyr	Leu	Ala	Leu	Glu	Asn	
15 Gly Lys Val Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala 115					100					105					110			
15 Gly Lys Val Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala 115																		
115   120   125   125   126   125   126   125   126		GGG	AAG	GTC	TTC	CTG	ACG	GGT	GGG	GAC	CTC	ccc	GCT	CTG	GAT	GGA	GCC	384
CGG GTG GAT TTC CGG TGT GAC CCT GAC TTC CAT CTT GTG GGC AGC TCC ATG VAI Asp Phe Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser 130  CGG AGT ATC TGT AGT CAG GGC CAG TGG AGC ACT CCC AAG CCC CAC TGC AAG Ser Ile Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys 145  CAG GTG AGC CGA ACG CCG CAC TCA GAG CGG CGA GCG GTG TAC ATC GGG Gln Val Ser Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly 165  GCG CTG TTT CCC ATG AGC GGG GGC GGC GGG GGC CAG GCC TGC CAG Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180  CCC GCG GTG GAG ATC GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC 180  CCC GCG GTG GAC ATC GGG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC 190  CCC GCG GAC TAC GAG CTC AAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC 191  CCC CCG GAC TAC GAG CTC AAG CTC ATC ACC CAC CAC AGC AGC AAG TGT GAC 191  CCC CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC CAC AGC AGC AAG TGT GAC 192  CCC CCG CAA GCT ACC AAG TAC CTC AAG CTC ATC CAC CAC CAC AGC AAG TGT GAC 195  CCC CCG CAA GCT ACC AAG TAC CTC AAG CTC ATC CAC CAC CAC AGC AAG TGT GAC 195  CCC CCG CCG CAA GCT ACC AAG TAC CTC ATC CAC CAC CAC AGC AAG TGT GAC 196  CCCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720	15	Gly	Lys	Val	Phe	Leu	Thr	Gly	Gly	Asp	Leu	Pro	Ala	Leu	Asp	Gly	Ala	
Arg   Val   Asp   Phe   Arg   Cys   Asp   Pro   Asp   Phe   His   Leu   Val   Gly   Ser   Ser				115					120					125			•	
Arg   Val   Asp   Phe   Arg   Cys   Asp   Pro   Asp   Phe   His   Leu   Val   Gly   Ser   Ser																		
20		CGG	GTG	GAT	TTC	CGG	TGT	GAC	CCT	GAC	TTC	CAT	CTT	GTG	GGC	AGC	TCC	432
CGG AGT ATC TGT AGT CAG GGC CAG TGG AGC ACT CCC AAG CCC CAC TGC ARG Ser Ile Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys 145		Arg	Val	Asp	Phe	Arg	Cys	Asp	Pro	Asp	Phe	His	Leu	Val	Gly	Ser	Ser	
Arg Ser Ile Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys 145	20		130					135					140					
Arg Ser Ile Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys 145																		
145 150 150 155 160  25  CAG GTG AGC CGA ACG CCG CAC TCA GAG CGG CGA GCG GTG TAC ATC GGG 528 GIn Val Ser Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly 175  30  GCG CTG TTT CCC ATG AGC GGG GGC TGG CCG GGG GGC CAG GCC TGC CAG Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180 185 190  CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC 624  Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile 195 200 205  CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC 672  Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720		CGG	AGT	ATC	TGT	AGT	CAG	GGC	CAG	TGG	AGC	ACT	CCC	AAG	CCC	CAC	TGC	480
CAG GTG AGC CGA ACG CCG CAC TCA GAG CGG CGA GCG GTG TAC ATC GGG GIN Val Ser Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly 175  GCG CTG TTT CCC ATG AGC GGG GGC TGG CGG GGG GGC CAG GCC TGC CAG Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180  CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC 195  CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC 195  CCC GCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC 195  CCG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC 672  Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720		Arg	Ser	Ile	Cys	Ser	Gln	Gly	Gln	Trp	Ser	Thr	Pro	Lys	Pro	His	Cys	
CAG GTG AGC CGA ACG CCG CAC TCA GAG CGG CGA GCG GTG TAC ATC GGG Gln Val Ser Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly 165		145					150					155					160	
Gln Val Ser Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly 175  30 GCG CTG TTT CCC ATG AGC GGG GGC TGG CCG GGG GGC CAG GCC TGC CAG Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180	25															٠		
GCG CTG TTT CCC ATG AGC GGG GGC TGG CCG GGG GGC CAG GCC TGC CAG Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180  CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile 195  CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720		CAG	GTG	AGC	CGA	ACG	CCG	CAC	TCA	GAG	CGG	CGA	GCG	GTG	TAC	ATC	GGG	528
GCG CTG TTT CCC ATG AGC GGG GGC TGG CCG GGG GGC CAG GCC TGC CAG Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180  CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC  Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile 195  CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720		Gln	Val	Ser	Arg	Thr	Pro	His	Ser	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	
Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180						165					170					175		
Ala Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln 180																		
CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC  Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile 195  CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720	30	GCG	CTG	TTT	CCC	ATG	AGC	GGG	GGC	TGG	CCG	GGG	GGC	CAG	GCC	TGC	CAG	576
CCC GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC  Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile  195  CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC  Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp  210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC  720		Ala	Leu	Phe	Pro	Met	Ser	Gly	Gly	Trp	Pro	Gly	Gly	Gln	Ala	Cys	Gln	
Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile 195  CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720					180					185					190			
Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile 195  CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720																		
CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210 CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720																		624
CTG CCG GAC TAC GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAC Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210 215 220  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720	35	Pro	Ala		Glu	Met	Ala	Leu		Asp	Val	Asn	Ser	Arg	Arg	Asp	Ile	
Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210 215 220  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720				195					200					205				
Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp 210 215 220  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720																		
40 210 215 220  CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720																		672
CCA GGC CAA GCT ACC AAG TAC CTG TAT GAA CTG CTC TAC AAC GAC CCC 720		Leu		Asp	Tyr	Glu	Leu		Leu	Ile	His	His		Ser	Lys	Суѕ	Asp	
	40		210					215					220					υ
											_							
Pro Gly Gln Ala Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro																		720
		Pro	Gly	Gln	Ala	Thr	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro	

	wo	99/2	1890							-						PCT/S	E98/01947
	225					230					235					240	_
	ATC	AAG	ATC	ATC	CTC	ATG	CCT	GGC	TGC	AGC	TCT	GTC	TCC	ACG	CTT	GTG	768
	Ile	Lys	Ile	Ile	Leu	Met	Pro	Gly	Суѕ	Ser	Ser	Val	Ser	Thr	Leu	Val	
5					245					250					255		
	GCT	GAG	GCT	GCC	AGG	ATG	TGG	AAC	CTC	ATT	GTG	CTC	TCC	TAT	GGT	TCC	816
	Ala	Glu	Ala	Ala	Arg	Met	Trp	Asn	Leu	Ile	Val	Leu	Ser	Tyr	Gly	Ser	•
10				260					265					270			
	AGC	TCA	CCA	GCT	CTG	TCC	AAC	CGG	CAG	CGC	TTT	ССТ	ACC	TTC	TTC	CGA	864
	Ser	Ser	Pro	Ala	Leu	Ser	Asn	Arg	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	•
			275					280					285				
15	ACT	CAT	ccc	TCG	GCC	ACG	CTC	CAC	AAC	ССТ	ACG	CGA	GTG	AAG	СТС	TTT	912
	Thr	His	Pro	Ser	Ala	Thr	Leu	His	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe	
		290					295					300					
	GAG	AAG	TGG	GGC	TGG	AGG	AAG	ATT	GCC	ACC	ATC	CAG	CAG	ACC	ACC	GAG	960
20	Glu	Lys	Trp	Gly	Trp	Arg	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	
	305					310					315					320	
	GTG	TTC	ACA	TCG	ACT	CTG	GAC	GAC	CTA	GAG	GAA	CGA	GTG	AAG	GAG	GCT	1008
	Val	Phe	Thr	Ser	Thr	Leu	Asp	Asp	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala	
25					325					330					335		
	GGG	ATT	GAG	ATT	ACT	TTC	CGC	CAG	AGC	TTC	TTC	TCA	GAT	CCT	GCC	GTG	1056
	Gly	Ile	Glu	Ile	Thr	Phe	Arg	Gln	Ser	Phe	Phe	Ser	Asp	Pro	Ala	Val	
30				340					345					350			
	CCT	GTC	AAG	AAC	CTC	AAG	CGC	CAG	GAT	GCC	CGA	ATC	ATC	GTG	GGA	CTT	1104
	Pro	Val	Lys	Asn	Leu	Lys	Arg	Gln	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	
			355					360					365				
35	TTC	TAT	GAG	ACT	GAA	GCC	CGG	AAA	GTG	TTC	TGT	GAG	GTA	TAC	AAG	GAG	1152
	Phe	Tyr	Glu	Thr	Glu	Ala	Arg	Lys	Val	Phe	Cys	Glu	Val	Tyr	Lys	Glu	
		370					375					380					
	CGG	CTC	TTT	GGG	AAG	AAG	TAT	GTG	TGG	TTC	CTC	ATT	GGG	TGG	тат	GCT	1200
40	Arg	Leu	Phe	Gly	Lys	Lys	Tyr	Val	Trp	Phe	Leu	Ile	Gly	Trp	Tyr	Ala	!
	385					390					395					400	
	GAC	AAT	TGG	TTC	AAG	ACC	TAC	GAC	ccc	TCC	ATC	AAC	TGC	ACA	GTG	gat	1248
	H18	65-1	wo	SEQ	• • • • • • • • • • • • • • • • • • • •			8	33								

	Asp	Asn	Trp	Phe	Lys	Thr	Tyr	Asp	Pro	Ser	Ile	As'n	Cvs	Thr	Val	Asp	
			•		405		•			410			-4-		415		
					,												
	GAG	ATG	ACC	GAG	GCT	GTG	GAA	GGC	CAC	ATC	ACC	ACT	GAG	ATT	GTC	ATG	1296
5	Glu	Met	Thr	Glu	Ala	Val	Glu	Gly	His	Ile	Thr	Thr	Glu	Ile	Va1	Met	
				420					425					430			
	CTG	AAC	CCA	GCC	AAC	ACC	CGC	AGC	ATC	TCC	AAC	ATG	ACA	TCC	CAG	GAG	1344
	Leu	Asn	Pro	Ala	Asn	Thr	Arg	Ser	Ile	Ser	Asn	Met	Thr	Ser	Gln	Glu	
10			435					440					445				
			_									CAC					1392
	Phe		Glu	Lys	Leu	Thr	_	Arg	Leu	Lys	Arg	His	Pro	Glu	Glu	Thr	
		450					455					460					
15	000	666	mm.c	030	030	CC3	000	OTTO	~~~	ma m	~ m	000		maa	~~~	mmo.	1.440
												GCC					1440
	465	GIY	Pile	GIII	GIU	470	PIO	reu	Ala	туг	475	Ala	iie	Trp	AIA		
	403					4,0					4/3					480	
20	GCA	TTG	GCC	CTG	AAC	AAG	ACA	тст	GGA	GGG	AGC	GGC	CGT	TCG	GGG	GTG	1488
												Gly		-			
					485	_			•	490			<b>J</b>		495		
	CGC	CTG	GAA	GAC	TTC	AAC	TAC	AAC	AAC	CAG	ACG	ATC	ACA	GAC	CAA	ATC	1536
25	Arg	Leu	Glu	Asp	Phe	Asn	Tyr	Asn	Asn	Gln	Thr	Ile	Thr	Asp	Gln	Ile	
				500					505					510		·	
	TAC	CGC	GCA	ATG	AAC	TCC	TCG	TCC	TTT	GAG	GGT	GTC	TCT	GGC	CAC	GTG	1584
	Tyr	Arg	Ala	Met	Asn	Ser	Ser	Ser	Phe	Glu	Gly	Val	Ser	Gly	His	Val	
30			515					520					525				
												ACT					1632
	Val		Asp	Ala	Ser	GIĀ		Arg	Met	Ala	Trp	Thr	Leu	Ile	GIu	Gin	
25		530					535					540				,	
35	רינולי	CAG	ദ്രസ	GGC	AGC	ጥልቦ	AAG	AAG	ልጥሮ	GGC	ጥልሮ	TAT	GAC	<b>AGC</b>	ACC	AAG	1680
												Tyr					1000
	545		~~ <i>3</i>	~~ 7		550	_, _	~, 0		1	555	- 7 *	٠٢			560	
	J <del>1</del> J					220					ددد						
40	GAT	GAC	CTT	TCC	TGG	TCT	AAA	ACG	GAC	AAA	TGG	АТТ	GTT	ACA	TCC	AGA	1728
												Ile					0
					565		•			570	. 4		,		575	-	
										_					-		

ACT CCC AGC CCA ACT TGA
Thr Pro Ser Pro Thr \*
580

1746

5

(2) INFORMATION FOR SEQ ID NO: 59:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 581 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

15

20

25

35

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 59:

Met Leu Leu Leu Leu Pro Leu Ala Leu Ala Pro Leu Phe Leu Arg

1 5 10 15

1

Pro Pro Gly Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly
20 25 30

Cys Gln Ile Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly
35 40 45

Leu Thr Arg Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr
50 55 60

Glu Ile Glu Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys
65 70 75 80

Val Arg Lys Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro 85 90 95

Ser Arg Cys Val Arg Ile Cys Ser Lys Ser Tyr Leu Ala Leu Glu Asn 100 105 110

Gly Lys Val Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala
40 115 120 125

Arg Val Asp Phe Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser 130 135 140

H1865-1 WO SEQ

	Arg 145	Ser	Ile	Cys	Ser	150	GIĀ	Gin	Trp	Ser	155	Pro	Lys	Pro	His	Cys 160
5	Gln	Val	Ser	Arg	Thr 165	Pro	His	Ser	Glu	Arg 170	Arg	Ala	Val	Tyr	Ile 175	Gly
10	Ala	Leu	Phe	Pro 180	Met	Ser	Gly	Gly	Trp 185	Pro	Gly	Gly	Gln	Ala 190	Cys	Gln
	Pro	Ala	Val 195	Glu	Met	Ala	Leu	Glu 200	Asp	Val	Asn	Ser	Arg 205	Arg	Asp	Ile
15	Leu	Pro 210	Asp	Tyr	Glu	Leu	Lys 215	Leu	Ile	His	His	Asp 220	Ser	Lys	Cys	Asp
	Pro 225	Gly	Gln	Ala	Thr	Lys 230	Tyr	Leu	Tyr	Glu	Leu 235	Leu	Tyr	Asn	Asp	Pro 240
20	Ile	Lys	Ile	Ile	Leu 245	Met	Pro	Gly	Cys	Ser 250	Ser	Val	Ser	Thr	Leu 255	Val
25	Ala	Glu	Ala	Ala 260	Arg	Met	Trp	Asn	Leu 265	Ile	Val	Leu	Ser	Туг 270	Gly	Ser
	Ser	Ser	Pro 275	Ala	Leu	Ser	Asn	Arg 280	Gln	Arg	Phe	Pro	Thr 285	Phe	Phe	Arg
30	Thr	His 290	Pro	Ser	Ala	Thr	Leu 295	His	Asn	Pro	Thr	Arg 300	Val	Lys	Leu	Phe
	Glu 305	Lys	Trp	Gly	Trp	Arg 310	Lys	Ile	Ala	Thr	Ile 315	Gln	Gln	Thr	Thr	Glu 320
35	Val	Phe	Thr	Ser	Thr 325	Leu	Asp	Asp	Leu	G1u 330	Glu	Arg	Val	Lys	Glu 335	Ala
40	Gly	Ile	Glu	Ile 340	Thr	Phe	Arg	Gln	Ser 345	Phe	Phe	Ser	Asp	Pro 350	Ala	Val
	Pro	Val	Lys 355	Asn	Leu	Lys	Arg	Gln 360	Asp	Ala	Arg	Ile	Ile 365	Val	Gly	Leu

Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Thr Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Ser Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Val Thr Ser Arg Thr Pro Ser Pro Thr 

((2) INFORMATION FOR SEQ ID NO: 60:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 16862 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

10

5

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

15 (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION: 3415..3440

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION:3441..3903

25

35

40

20

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION: 3904..3988

30 (ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION: 3989..4689

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION: 4690..4893

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION: 4894..5645

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION:5646..5831

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION:5832..7181

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION: 7182..7202

10 (ix) FEATURE:

5

20

30

40

(A) NAME/KEY: intron

(B) LOCATION: 7203..8307

15 (ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION:8308..8803

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION: 8804..12266

(ix) FEATURE:

(A) NAME/KEY: exon

25 (B) LOCATION: 12267..12401

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION: 12402..12815

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION:12816..12986

35 (ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION: 12987..14085

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION: 14086..14187

(ix) FEATURE:

PCT/SE98/01947 WO 99/21890

(A) NAME/KEY: intron

(B) LOCATION: 14188..14473

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION: 14474..14539

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION: 14540..14998

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION: 14999...15190

15

10

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION: 15191..16862

20

30

35

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 60:

GATCATATTA ATTTGAAGGT GGCGGGGCAG GATGGTTCTG TGGTGCAGTT TAAGATTAAG 25 AGGCATACAC CACTTAGTAA ACTAATGAAA GCCTATTGTG AACGACAGGG ATTGTCAATG 120 AGGCAGATCA GATTCCGATT CGACGGGCAA CCAATGAAAC AGACACCT GCACAGTTGG 180 AAATGGAGGA TGAAGATACA ATTGATGTGT TCCAACAGCA GACGGGAGGT GTCTACTGAA 240 AAGGGAACCT GCTTCTTTAC TCCAGAACTC TGTTCTTTAA AGACCAAGAT TACATTCTCA 300 ATTAGAAAAC TGCAATTTGC TTCCACCACA TCCTGACTAC TACCGTATAG TTTTCTCTAT 360 TCTTTCATTT CCCCCTTCCC CATTCCTTTA CTGTACATAA AGTAACTGGT ATATGTGCAC 420 480 AAGCATATTA CTTTTTTTT TTAAAACTAA ACAGCCAATG GTATGTTTTG ATTGACATCA AGTGGAGACG GGGGGAAAA TACTGATTCT GTGAAAATAC CCCCTTTCTC CATTAGTGGC 540 ATGCTCATTC AGCTCTTATC TTTATATTCC AGTAAGTTAT TTTGCTCTCA CTGTTTTAAC 600 AACAACAACA AAAAAACAAC AACATAAAAA TCCTTGCATA CCTTGTTCAA TTGGAGAATT 660

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	TTAATGTTTT	TCATTTATCA	TTGTAAAACC	AAGGACAATT	TTATAACTTT	TTTGTACTTA	720
	GCTGTTACAT	GCAGAGCAAT	CTGTCTTTAA	GTAGGGATAA	АТТАСТСТАА	AACAAAAAAG	780
5	AATCCTAGAT	AGTTTTCCCT	TCAAGTCAAG	CGTCTTGTTG	ТТТАААТААА	CTTCTTGTTT	840
	AAAAAAAAA	AAAGTAAAAA	AGAAAAGTTA	TGCAACAATT	AATGGCCCAG	AGGCAATCCT	900
10	TGTTAACATT	TTGATGCATC	TTTTAGCTGT	TTTTTTTTT	TTTTTTTT	TTGACTGAGT	960
	TTGACTCTTG	TCACCCAGGC	TGAAGTGCAA	TGGCATGGCA	TGATCTTGGC	TCACTGCAAC	1020
15	CTCCGCCTCC	CGGGTTCAAG	TGATTCTCCT	GCCTCAGCCT	CCTGAGTAGC	TAGGATTACG	1080
13	GGCATGCACC	ACCATGCCTG	GCTAATTTTG	TATTTTAGT	AGAGTTGGGG	CTTCTCCACA	1140
	CTGGTCAGGC	TGGTCTCGAA	CTCCCAACCT	CAGGTGATAA	GGGAAGGGGC	ACTATTGACA	1200
20	TTTATGGTTG	GGGCAGAGGT	GTAAGATATT	CTTCAAAGCA	CTACCTACAT	GTTGAAGAAT	1260
	TGTTCCTCAC	CCAGATTCTC	AAAAGTCCCC	CAGGACATTC	ACGTAGTGAA	AACCTGTGTT	1320
25	TAATTATCTG	AGCCTATAAC	TTAATACAGT	TTTAAAATTT	TTTTTTAAAT	ATACAGTGAA	1380
	CTTTCTAGGA	ATGCAATTAT	AGTTGTGTGT	AAAATTAGGG	AAAATTAACT	TTGCTACCAA	1440
	GAGTTGTTCA	ACATTTTGTT	AAATCACTTC	ATTGATGGCA	ACATGCTGGA	GGTAGTTGAG	1500
30	TCACCAACTC	AGCACCTGGA	TCAGCCTGTG	TTGGTAGCAG	TTTCATCCCC	GTGGTTCTGT	1560
	GAATAGGTGG	AAGCATCTGC	TTACTCCATC	AGGACTTCTA	GGGTAGTCGG	GCCTTGGCAC	1620
35	TCACACATTA	AAATACTGTT	TATGTTATTT	TATTGCAAGT	TACTTTTCTT	TCATTTCCCC	1680
	TTTACGTTAC	AGAAAGGGAA	GCATTTTGCT	TTCTGTTTAA	AGTTGTGTAT	GTAGGTAGGT	1740
	TATATCATCT	AWGACTTTCT	CTCCCTCCTT	CCCTTTCTTT	TTGTTTGAGA	TGGAGTCTTG	1800
40	CTCTGTCACC	CAGGCTGGAG	TGCAGTGGTG	CGATCTTGGC	TCACTGCAAC	CTCTGCCTCC	1860
	CGGGTTCAAG	CGATTCTGGT	GTCTCAGCTG	GGATTACAGG	CGCACACCAT	CACACCACGC	1920

	TAATTTTTCT	ATTTTTAGTA	GAGATGGGGT	TTCGCCATGC	TGGCCAGGCC	AGGCTGGTCT	1980
	CAAACTCCTG	AGCTCAAGTG	ATCAGTCCGC	CTCGGCCTCC	CAAAGTTCTG	GGATTTCAGG	2040
5	CGTGAGCCTC	ATCTATGAAT	CTCAATTTAG	GACAGTAAAA	GTGTCATTAC	АААААТАТТТ	2100
	ATTGTAAAAA	AGGGTTGGAG	GTTGAGAATC	TCAATTCTAG	TCAGTCTCTC	AGTGTTTGGT	2160
	TTCTTCCTAC	CATTTTTCCC	CCTAGGACCA	GCCAGAAAGC	AGCTTTTTT	TTGTCCCCCC	2220
10	CAACAAGGAG	CCCACTGTTT	CCTCTCCCAG	CCCAAACTCA	GGCCTACGAA	CAACAACAGC	2280
	ACTACACACA	CACACACACA	CACACACACA	CACACACACA	CACCCCTCCA	CTTCAAGGTA	2340
15	TAGCCAAGAG	CTTCTGGAGC	CGTCAAAAAG	GTCTGTACCT	GCTGTCTTTA	GAGCTTCCAG	2400
	TTTGCCCTTG	GTCAAGAAAT	ACTGTTTGCT	AGGCTCTGCT	GGAGTACATC	AGGTAATACT	2460
	GGCTTCTAAA	CCACCCTGAG	GTTCTTTTCT	CTTGTCCTTT	TACTCCCTTC	GTACTTCAAT	2520
20	TTCTCTCCTT	GATGTCCCCC	TCCCTGTTTT	GTTTTTTGCC	TCCAATCCGT	TCTGCGCGTT	2580
	CCCTGCAGAG	CAGGCGAGTA	GCAATGCTGC	TGGACCATGG	AGCTGCTCTA	GTCTCCCAGA	2640
25	AATCTCTTCT	ACACCCAACC	CTTCTTGCGC	TTAGGTGGTC	CTCAGTCCCC	CTCCCCCACC	2700
	TCCTTCTGAC	CCAGGCTTCT	TTCTCGCCCT	CCGGTCGCAG	TTCTCCTGGG	CATCTGCCTC	2760
30	TGCCTCTCTC	CTCTCACCCG	GATCTAGGGC	TGCCTTCTCT	TTGTGCAGCC	GTCTTTCTCC	2820
30	ACCTTCATCC	CAGACTCCCT	GTCTCAGCGC	CAGCTCCTCT	GCCTTTGGCT	CGGGTTCCCT	2880
	CTCCCCCACC	CCAGCTTCCA	GTTGTTTGGC	CCGCAGGTCC	CTCGGCAGTG	ACCGGCGCCC	2940
35	CCCGACGAGT	GCGTGTGCAC	CAGGGCACCT	CCCTCTCCCC	CACCTCTCAG	CCCCGCGCCT	3000
	CTCCACCGCC	CGCCCCACCG	CGCTGTGGGC	GGTCCAGGGC	GGGGCTGGGA	TCCGGGGCGG	3060
46	CTCCCGGGGC	TCGGGTTGTG	GGAGGCGCCC	TCTCCCCGGT	CTTCCCCTCT	CTTCCCCCCG	. 3120
40	CCCTGCCTTC	: CCTTGCACCC	: TCCTTCTTCC	CTCCGCCCGG	GAGCTCTCCC	TGGTCCCCGG	3180
	CGCCGCCTCC	TTCCCTCCC	GCTCCCCGCI	CCCCGCTCCC	GTGGCTGCCG	CCGCCCCGGG	3240

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	GAAGAAGAGA	CAGGGGTGGG	GTTTGGGGGA	AGCGAGAGAG	GAGGGGAGAG	ACCCTGGCCA	3300
_	GGCTGGAGCC	TGGATTCGAG	GGGAGGAGGG	ACGGGAGGAG	GAGAAAGGTG	GAGGAGAAGG	3360
5	GAGGGGGGAG	CGGGGAGGAG	CGGCCGGGCC	TGGGGCCTTG	AGGCCCGGGG	AGAGCCGGGG	3420
	AGCCGGGCCC	GCGCGCCGAG	GTAAGAGCCA	GGGCCCCGGG	TTAGCAGGGC	TCGGAGAGGG	3480
10	GGCGCGCGC	GTGGTGGGGG	AGGGGGCAGT	GGGCGCAGGG	CCCAGCTGGG	GGAAGCGGGG	3540
	CTGGGGGAGA	GGAGGAACCG	CGGGGATGGA	ATCGGGGAGC	GCTGAGGCGG	CCGATGCCGG	3600
	GAGCGTGGGT	AAGCCAGGCT	TCTGCGAGCC	GCGGGGGCCG	GGGGAGAGGA	GGTGGTGAGA	3660
15	GGTGGAGTCC	GGGAGGGTTG	GGGGCCGAGG	GAGGCAGGAG	GAGGGTGGGG	ACAGGCTTTC	3720
	TCTCCTCCTC	TCCCCCACC	CCGCGCGGGG	CTCCGCCCCC	GCCTCCTCCG	CGGGGCGCTC	3780
20	TCTTGGTCCC	CAGGCTGAGC	CCGGTCGGAG	CCTGCGAGGC	AACCGGCAAG	AGGTCGAGTA	3840
	GTCTCCGGGT	GCGGGCCGCG	CCGGCGGGGC	TCGGTCCAGT	CCTCATGGCC	GCCTCTCACT	3900
25	TAGATGTTGC	TGCTGCTGCT	ACTGGCGCCA	CTCTTCCTCC	GCCCCCGGG	CGCGGGCGGG	3960
۵	GCGCAGACCC	CCAACGCCAC	CTCAGAAGGT	GCATCCTTCT	TCGACGACCT	CCGGCCCTCC	4020
	TTCGCTCCAC	TTCCCTTTCC	CTGCATCTCC	TCATTTCTGG	TCCTCATCAC	TATCCCATCA	4080
30	GTCCCACATA	TCATCCCGGT	CTGGCAACCC	CTTCTGCTCG	GCCCGACTTT	ACTACTGCTG	4140
	ACCTCCTTCT	GTCACCCCAC	GTTACTATCC	AGCACCTCTT	TTCTCTGCCC	ACATTGCTAC	4200
35	ACTATACCAC	CTTCCTGTGC	ATTTTCTCCG	CCTCAATCCC	CTTTCCCAGC	CCCACATTAC	4260
33	TACCTCAATT	ACTCCCTTTT	CTTGGTCCCA	CTTTGCTGTC	CAGATGATCT	TATTAGCCTC	4320
	CCTTTATCCT	ССТАТССТАА	TTCAACTGGA	ATATCCTCAT	TTAGCCTTTT	TTTTTAAAGA	4380
40	AAAGCTCCAC	CCACATATCA	TACCCTTCAT	GATTTCTTAA	TTACTTTTCT	TTCTTACCTC	4440
	CACCCAGCAC	CCTTCCCTCC	CCACTTGTGG	GTTCTCTCAT	CAGCTTTAAC	CCTGGCCCTT	4500

	TACTCTCTGT	CCTTTAGCCA	GGGGATCTGT	ACCTGTCCCC	ACTCCCACCC	TCTAGTGCCC	45.60
	CATCCCTCTT	CCTCTGTCCC	CAGCCTGCCC	ACAGACCACG	CCCTACTCTC	CCCTTCCTCC	4620
5	CACTGGGGAG	CCTGCCTTTT	CCTCTTTCCC	ACCATTCCTC	TCTGTATGCC	TCCCCGACTC	4680
	ACCCCTTAGG	TTGCCAGATC	ATACACCCGC	CCTGGGAAGG	GGGCATCAGG	TACCGGGGCC	4740
	TGACTCGGGA	CCAGGTGAAG	GCTATCAACT	TCCTGCCAGT	GGACTATGAG	ATTGAGTATG	4800
10	TGTGCCGGGG	GGAGCGCGAG	GTGGTGGGC	CCAAGGTCCG	CAAGTGCCTG	GCCAACGGCT	4860
	CCTGGACAGA	TATGGACACA	CCCAGCCGCT	GTGGTGAGTA	GCCTCGGAAG	CCCCTCCCCT	4920
15	CTTCAAGACT	ATTCCTTTTC	CTGCCGCAAA	CTTAGCATTA	CTGCTTGCAA	GTCAGCACTT	4980
	TAAATCCAGT	ATACCAAAAT	TCACAAATAC	ATTTATTGAA	TGACTACTAC	ATAAGAGCAA	5040
20	TTTTGCTCTG	TGCGGTTGGA	GGTAGTAGAG	CTAGCAGCCT	GCACAGTTCA	TTTCATCCTC	5100
20	CCTTCATTAG	GCCACTGATC	ATTGGCCTAT	AACATTGATA	ATTCATCTTG	TCAGTTATTC	5160
	TCTTTGAGGA	TCATTAGTGG	CAGATGATGA	CAAAAAAATT	CTAAAATGAT	TTCATCACAT	5220
25	TTTTGAATAC	CTCTGTCACC	AACCCAGAGA	CCATATGCCC	AAGAAACAAA	AGCCAGTTTA	5280
	ATATTAATAG	AAGCCAACTA	TAATAAGAAA	AGCAAATCTG	ATTGTGCATC	CAAAGTTATA	5340
30	TACATCTACA	TATTTCAAAG	CCAGAGAACC	GCCCACTGTA	GCTGACTTTG	AAGAGATCCC	5400
30	ATTTTGTGTG	CTTATAGCCC	CATCTTGGGT	TCCTAAAATG	GTAATTTTTT	TTTTCTTTTG	5460
	GGAATGTGTG	GATGCTTGCA	CAGGTAAGGG	AGGATTGGAA	GATAGGTAGG	CAAATCCTTT	5520
35	TCACATGTGA	TTTTCTTTAG	AGCAGGATGC	TTGTGGACCC	AAACCTGCAC	CTGAGTCCCC	5580
	TGCTCTTTAA	AGGGAAAGAG	CCTTCTTCAA	CTCGCCTCTC	TTCTTATTTT	CCTATCTCTC	5640
	CACAGTCCGA	ATCTGCTCCA	AGTCTTATTT	GACCCTGGAA	AATGGGAAGG	TTTTCCTGAC	5700
40	GGGTGGGGAC	CTCCCAGCTC	TGGACGGAGC	CCGGGTGGAT	TTCCGGTGTG	ACCCCGACTT	5760
	CCATCTGGTG	GGCAGCTCCC	GGAGCATCTG	TAGTCAGGGC	CAGTGGAGCA	CCCCCAAGCC	5820

	CCACTGCCAG	GGTGAGGGGA	ACAGCTGCCT	GCATGCAGCT	GATGAGGACG	CTTGTGTGAG	5880
5	GATGGGAGTG	GGGTGGGAAT	GGATAATGGG	AAAGAATGGA	GAGCTATAAA	AATGTGGGGG	5940
	AGGACACTGG	AAAGGGGAGA	TGAAAGTCCC	TTTTTCCTCC	ATCACCTGCC	TCAAACTTCC	6000
	TCTTGCAGTC	CCCGGTATCC	TCTGTAGGTT	GGGGGCTTCC	TTCCTTTACC	TTTTAAAAAA	6060
10	ATCTTCCTGC	TCCCGATTCT	TAGACCTCAC	GTTTTCTCTT	TTCCTTTATG	AATCTCACCT	6120
	CTCTCACCTT	CTTCAGGTTT	AAATACTCCA	ATTTTCCCTT	TCTCTAAACT	TAGAAATTTC	6180
15	CATGCATCAC	CCTCTTCTAG	AATTCATCCC	TCACCATTCC	ТТАТАТАТТ	GATTTATTGT	6240
	AAAGACTCAG	AAATAAATCA	AACATTCTAC	TAAGAAAAAT	TGAGAAGGGG	AGCTCTGGGG	6300
	GTGGAAACAT	ATTAGGGTAA	AAGACTTAAA	ATTGGAGGCA	GCATTATCAG	AAGATGAAGA	6360
20	ACAACTCAGG	GATGGGGTGG	GAAGAAGACA	GGTCCTTTTC	TGKACTTCCT	AGACAACCTC	6420
	CATTATTCCC	TAAGGGAATC	AGTGTTGTGT	CTGTCTACYT	TTTTTTTT	TTTŢTTGCCA	6480
25	CGTAATTTTA	CAAACTCTCC	CTTTTCTAGG	CACCCGAACT	CTCTGCCATC	TTCTCTCCTG	-6540
	GGATGCAGTC	ATCCCATTTG	TATGCCTCAT	ACTTCCTCTA	CCCTGGTAGA	TTCTTTCAAG	6600
	ATCCTTGGGC	TTTACTTTCC	TCACATAACT	CAGTTATTCT	GCTTCTAGTT	ТАССАТТТТА	6660
30	TTCTGGAAAT	TGAGAGTCCC	ATCCAGGGGT	GGACTTATGA	CACTACTGAA	ACTTAGACTT	6720
	CAAGGTTCCT	CACCTACAGG	GCCCTCTTCC	TGTGCTCTAA	TAATATAGAG	GGCTCGATGG	6780
35	ATATGTGTTC	ATATGGTAAC	AGGCTTTTGT	AAAAATTGCA	GAAATAAGAT	TTTAACAGCA	6840
	ATTGCTTAAA	GCCAATTGTA	TGTGTAATTT	TTTTTCTTAA	AGACTCCCAA	TTTTGTAATA	6900
	TTCAGGCACC	ACAGAACCAA	GATCTGCCCC	AAACTTAGCT	ATTGGCATTC	CCGTCTCAAA	6960
40	TTCTGTTGTC	CTATGAAAAA	TCGAAGAAGA	AAATAAGTCC	TGACCCCCTT	ACCCCCAGAC	7020 u
	CCACCTTGTT	CTTATCCCCA	GGCACCCTCC	CCTCAGAAAC	GCAGGCTTCT	GCTCTCCCCG	7080

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	GTCTTCAGCA	TGGACAGGTG	TGGGAGGGG	CTGGGGATCA	GGCCAĠGGAA	GCTGGGCGCC	7140
	AGTGGTAACT	CTTCTCTGAT	CCCCGTCTTT	CCTGCTGCCA	GTGAATCGAA	CGCCACACTC	7200
5	AGGTGAGATG	AGAAACCCTT	ACCGCGCGCA	CTGCAATGCC	CTCCCCTTCA	CTCTGCACCC	7260
	TCCACCCCC	TGAAATTCTG	CCCTTAGGCT	ACGGGGCGTC	GTCCTTTCGC	ACCTTCCCCA	7320
10	ACCCACCCCA	GTTTGCGGCC	ACCCCCTTCC	CTCCCTACCT	GTTTCCTGCC	TCCAGTCCCG	7380
10	GTTTTCCACG	AGGCTGCGGT	CTCTCCTTGT	CCCTGCTTGG	CTACACTTCC	CTGGGCTCCA	7440
	CCTCCTCCCA	GACTGAGCCT	CGCCGGTGTC	AGGCAGAGCC	CAGCAGARGG	CGGCAGGGTG	7500
15	CTGGGAGACC	CTGAGCTCCC	ACCACGTTTT	CCCCTGTGGG	GTTCCTTGCG	ACCTTCGCTG	7560
	GAACCTTTTC	CAGCCTGCTG	CCTCCTAGGA	TTTCACCTAA	TGGACTTTCT	CAGCCTGTCC	7620
20	CACCCATCCC	AACCCTGGCC	AGGCCTCTCG	CGCTCTTCCC	CACATCTTTT	CCTTCCGTGT	7680
20	ACCCCTTCCC	TCGTCTTTTC	TCAATTCCAT	GTCCTGTCTC	CCTTTCTTAG	GCTTCTGTCT	7740
	ACCCAGCCCC	AGGCTCCCTT	CCACGACCCC	ACCACTCCCT	CAAACCAGCC	TCCCTTCCGT	7800
25	ACCCAACTCG	TTCCCTCCAA	AACCGTTTCC	TCTCCCCCAC	ATCCTCAGTG	CTTCACTGTA	7860
	TCGACTCATA	CTCCCACTTC	AGACCTCAGG	CGCCAGCCCC	GTTTCTCTCC	CGTCCCACTC	7920
30	GCATCCTTCC	CTTCCTACCC	TGGTTCCTCC	GTGCTTCAGC	CTCCCGCGGC	TCCCTCCGCC	7980
	CACCCCCCCC	TCCTGGCACG	CCCCGTCCCC	ATTTCTCCTC	CCCTCGGGTC	CCCTTAAGTG	8040
	AGATCCCTCC	CTTCCTCTTT	CGTTCCTTTC	CTCCTCGAGG	TTGCATCCCC	сстсссстсс	8100
35	CCGCCCCTCC	GACTGTCGCT	CCCACCTCGG	CGCTCGCTTC	CCTCCCCGCC	CCCTTCCTGC	8160
	CTCCCCAGCT	cccgcccgcc	CCCCACCC	CCGCTGCCGC	GCGCCGCCG	TGACGTCAGA	8220
40	GCCCCTCCC	AGCCCCACAT	CTCCCTCCTG	СТССТССТСС	TCCCCTCCGT	CGGTCAGTCA	. 8280
••	GTCCGCGAGG	AGAGTCCGCG	GTGGCGGCGA	CGGTGGCGAG	AGCCGCGGGG	GCCGTAGGAA	8340
	GCCAACCTTC	CCTGCTTCTC	CGGGGCCCTC	GCCCCCTCCT	CCCCACAAAA	TCAGGGATGG	8400

	AGGCGCCTCC	CCGGCACCCT	CTTAGCAGCC	CTCCCGGGA	AAAGTGTCCC	CCCTGAGCTC	8460
•	CTAACGCTCC	CCAACAGCTA	CCCCTGCCCC	CCACGCCATG	GGGCCCGGGG	CCCCTTTTGC	8520
5	CCGGGTGGGG	TGGCCACTGC	CGCTTCTGGT	TGTGATGGCG	GCAGGGGTGG	CTCCGGTGTG	8580
	GGCCTCCCAC	TCCCCCCATC	TCCCGCGGCC	TCACTCGCGG	GTCCCCCCGC	ACCCCTCCTC	8640
10	AGAACGGCGC	GCAGTGTACA	TCGGGGCACT	GTTTCCCATG	AGCGGGGGCT	GGCCAGGGG	8700
	CCAGGCCTGC	CAGCCCGCGG	TGGAGATGGC	GCTGGAGGAC	GTGAATAGCC	GCAGGGACAT	8760
15	CCTGCCGGAC	TATGAGCTCA	AGCTCATCCA	CCACGACAGC	AAGGTAGCCC	TGGACATGGG	8820
15	GGTGGGTGGG	AGGTGGGGGC	TTGCGGGSCA	GGGGCCAAG	CAAGCCTGCA	CGCGCCCCCA	8880
	TCTGTCTGAG	TCGTCTCTGG	GATTGCGAGG	CAGACCCCTC	CCTTGTGTGA	CTGGCAGGAG	8940
20	ATGGGCTGGG	GGTGCAGGAG	CTTGGGAAGA	GTCGCAGGGG	CTGGAGGTCC	AAGATGAGGG	9000
	TCTAGGGGCT	CAAGATGGTT	AAGCATGCTG	CAAGGCAGAC	CCTTCTGCCC	CGCTGCGGGA	9060
25	GTCTCGCAGA	AGTGTCGGGG	TTTGGAGAAA	CTGGTGGTGG	ATTTAAGGTA	TTAGGAGACA	9120
•	CTGATCCTCT	GAGGGAGTAA	ACTAACCCTG	GAATGGGTTG	GGGGTGGAGG	GAATGTCAGA	9180
	GGTGGGGAGC	TGGATTGGGG	GGTTACATTT	ACCATGGTAA	CAAGGTAAAA	TCTTGGCGTA	9240
30	GGTTGGAGCT	GGAAGGAATA	GGGACAGAAT	GAGGAAAATT	TTGAGAGACT	TGAGAGCTCT	9300
	AGTTTATTTA	TCTTAACAAA	ACAGCAAGGT	AGTGGTGAGC	CCTACCTGAC	TCCTTCTCAT	9360
35	CCTTCTATTC	CCAACCCTGT	TGAGCATTCC	CAGACTGTGG	GATAGATGGC	ATATGGTGAT	9420
55	TGGGGAAGGC	TAATGATCAA	GAGGTGGGCA	GAGGCACTGG	GAAAATGAAT	TGGATTGGGG	9480
	ATCCACATGG	GAACCCCCAC	AATAGCATGG	GGATGAAGAA	GAGTCAACAT	ACAAGGAGAA	9540
40	GAGAACAGAA	AAGAATGGCA	GTGGGGGAGA	GGGGCAAGGA	GGTAGCGTGG	GGATAATGAG	9600,
	AGATCTTGGG	GCACCTTATG	GAACTTGGGT	CCTGACCTTC	CCTTCCCTTA	<b>TAGCATTGTG</b>	9660

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	GCCTCTAGGA	TGTGAGAAGG	GAAATGGGAT	GTAGGGATTA	GGGAGGTGAG	TTGÄGGGÄGA	9 <del>72</del> 0
	GAGAGAAGGT	AAGCAAATTT	GGGTCCAGGG	GTATTAGGGG	ATAGCTTATA	ATGAGGTTTT	9780
5	TTTTCCCACC	CCTCTCCCCT	ACATGAATAA	TTGGGGGTGC	AGGGAAGGAT	GTGACACAGG	9840
	GAAGGAGATT	TAAGATCTCA	AATTTATCTT	CACTGACATG	TGGCCCCAGA	GACTTAAGGA	9900
	ATTGGGTTAG	GGTGAAATAG	AGTACACAAG	GTGAGAATTT	GGTGATCTTA	CCAAATATCA	9960
10	ACCTTGGGGT	GATCCAAGGA	ТТТАТАТТСА	TTTTTAGAAC	ATCACTATAC	ACCTAGAAAT	10020
	AGGTGTGTGT	CTGGGATAGG	TGTGTGAGGG	GACAGAAGTG	AGGTTGAAGG	TAGGGTGCTT	10080
15	GAAGAGAAGA	GAGCACAAGG	ATTATCAGGA	GCTTGGCAAG	AGAACTTAAA	ATCCTTTTTG	10140
	ACTGTTACTT	TCTCGTGGTT	CTCAGCCTTC	AGTGTACATA	AGAATCACCA	GAGGAGTTTG	10200
	TTAAAAATAC	AGATTCTAGC	TCCTTGGTCA	GGGATGAATC	CCAAGTATTT	ATCTGTATTT	10260
20	TTACTAATAG	ACATCCTATC	TTGGTGGATT	CCTGAGCTGT	AAGCTAACCC	CAGAATGCCT	10320
	ATGGGAAGAG	CAGCAGGGTA	CAGGAAAATA	ATTAGGTATT	AGGGTACGGG	AGGCAGGAAG	10380
25	AGAAGTAGAG	GATCAGATCT	GGTAGAGGGT	CAGACTTGGG	ACAGTCAGAG	AGATCATTGG	. 10440
	TTTTGGGGAG	TGGAGTGTGA	AGAAAATGAC	AGGGAGAGAT	GGGTGCAGGC	TTTATGATAG	10500
30	GGGATCACAG	GAGATAGGGG	AGGCCTGGCT	GTGAGCTCAA	ACTCATCCAC	CATGACAGGT	10560
30	GATTCCCTGG	AGGTGGCGGG	GAGCAGACGT	GGGACCTGGG	AGAAGGGAAC	TGGAGAACAT	10620
	CAGAGGCATC	AAGCGGGGTG	GGATGGGAAG	GCAGAAGAAC	CAGAATGTGT	CAATTGGAAT	10680
35	GAGTCGGTTT	CCTGCCTGCA	AATCCAGATC	CTTGCAAGAG	CAAAGAGAGG	GAGGAGAACT	10740
	AAGGAAATCT	ATTGGGGAGG	GGGAGAGAAT	CACGTGGTGG	AGAGAATCTG	CAGTGATGAA	10800
40	TAGTGTGTGG	.AAGAGGGAAA	CGGTTGCAAG	AAAAGGTAGA	TAAGAAATCA	GGAAACAAAA	
40	TGGGGGCAT	GCCTTGCCCT	GTTGATATGT	ATCTTATATG	TTCTTGAATG	TCCTCATTGK	10920
	TCCTATTAAC	CCTGTCTTA	GAGAAGTGGA	GGGGCACTGA	GGGGCTGTGG	GAGAAGCTGG	10980

	GAGCAGGATC	TGGAGTAATA	GATGTGGGGA	GAGTGCAGGA	AGGTGGGTCC	TGAGAATGGT	11040
	AAAGATTTAC	AAAGTTGCCC	TAGTGGGAGG	CATAAAGAGA	AAACCTTCCA	ATGTTGTTGA	11100
5	GCACTGCCCT	TGGCCAGAGT	GAGGGTAGGG	TGGGCAACAG	AGAATTCTCA	GTGACTGCTG	11160
	GTTCTTCAGA	TTCCAACAGC	TTCCCCTGGC	TCCCCCTTCT	CCAACTTCCC	ACCGTGTCCC	11220
10	AAATGTCAGG	CCTCAGTGGG	AGGTAAGCAG	GCTCCAGAGT	GCTTTCTTTA	TTTCCTTTCT	11280
	ACTTATCCTC	·CCCTCCTGGC	AACATTTCAC	CCTCCTTAGT	CCCCTGAGCC	CCCTGTCTGT	11340
15	GTCCCCTCTG	CCCTGGCTCC	CCACTGGCTG	CCATTTCGTC	TTCACATGCA	TTGGGGTTCC	11400
13	AGCAGCTTCT	GAAATGTCAT	ATATCAGTGG	GAGGGGAACA	GGCAGTGGGA	GACCCAAGGC	11460
	TGGCTCTTCC	TCCCCCATTT	CCCCTCCTCC	CAAGCTTCCT	ТТСТТСТССА	GCTTTCTGCT	11520
20	TGTTTACTTT	CCCTAGCTCC	AAGCCTCTCT	TTAAGGCACC	TCTCAAATTG	KCTGGTTTCT	11580
	TGAGAGTTCC	ATTCTATTCA	TTCTCTCTGT	TCTTTCCTCA	TCCTACATTC	TTCCCTACTT	11640
25	CCACCCCCA	GTGTCTTTTT	TTCTAATGGA	CCTGTCAAAT	GTCAGCGCCC	AGCAGGAGGG	11700
23	ATGGATCACT	GAGCGGGACC	CCCTACTGGT	CTTGTTCCTG	TTCTCTCTTT	ACTTATCACT	11760
	AGCTCTGAAA	AGAGAAGAGG	GAGGAAACAA	ATGGAAGGTG	GGGAGAAGGG	GTTTGCAGAG	11820
30	GTGAGGAAGG	ААТТТТСАТА	ATATGGCTTT	GAGCAAGCTA	TCTGGGGATG	TGGAAAGAGT	11880
	TTACCGTATT	CCTACTGACT	TCTTCCACCC	ACTGGTGTTT	GAAGCATAGA	AACATGGGGT	11940
35	AAAGGGCTTG	GTGACAGAGG	GAAGGGGGAT	GTCTGAGGGT	GAGCTGAAAG	GAGGTAAGGT	12000
33	GGTATGTTCA	ТТААТАССАА	AGGAGGGGTG	TGCAGGAGAG	GTGATGGGTA	AGGCTCCAGA	12060
	TGGAAGACAG	AGAAGGAAGT	TTAATGAAAG	AGGAGAAAA	AGGCACTTGA	CAGGAAGAGA	12120
40	TGCCAGAAAG	GAGAAGAAAA	CGGTAATTAA	TGATGAAAGT	GAGTAATTGA	GAAAGGAACT	12180
	AATTTGTTCG	AGAAAGATAA	GAGCAGGAAT	TGCAGACAGG	GGAGGGCCC	CAGGAGAGCT	12240

	TGCCCTCATC	TCCTCTTGTC	TTTCAGTGTG	ATCCAGGCCA	AGCCACCAAG	TACCTATATG	12300
	AGCTGCTCTA	CAACGACCCT	ATCAAGATCA	TCCTTATGCC	TGGCTGCAGC	TCTGTCTCCA	12360
5	CGCTGGTGGC	TGAGGCTGCT	AGGATGTGGA	ACCTCATTGT	GGTAAGCAGG	GCTATGGGGG	12420
	TCAGAAGATG	GGGTCATTCC	CTTTTGAGCT	CTACTGAAGG	GACGATGGCG	ATTGTGGGTT	12480
10	TGTATTGAAA	AGGAGTGTGG	AGGACCTGCT	ACTAAGATTC	AGAGTCCTCT	GCAGACCTGA	12540
10	GCTAGGCAGC	CTCCTAGCAA	CAGTGGCCTG	ACAGTGCTGC	AGCTGACCTC	CTTCTTCAGA	12600
	AGGAATTGAA	ATTAGATCAG	TGAAAGAGCA	TCCCGGTTGT	GAGGGGTGTG	TGGGCCTTTG	12660
15	AGAATCTCTT	TTCCTTAGGC	AGACCAGAGG	TGGGGAGGTT	TGGAGAGAGT	AAGGAAGAGA	12720
	AACCCAAAGG	CAGGAAGAGG	GTTAAAGGAA	CTCTTGGCCA	CTCTTGGTGT	CCTCAGTGAA	12780
20	CAGACCCTGT	TGCACTCACT	CTCCCTGCCC	CACAGCTTTC	CTATGGCTCC	AGCTCACCAG	12840
	CCCTGTCAAA	CCGGCAGCGT	TTCCCCACTT	TCTTCCGAAC	GCACCCATCA	GCCACACTCC	12900
	ACAACCCTAC	CCGCGTGAAA	CTCTTTGAAA	AGTGGGGCTG	GAAGAAGATT	GCTACCATCC	12960
25	AGCAGACCAC	TGAGGTCTTC	ACTTCGGTGA	GGAGGGGTTG	GGCAAGGGGT	AAAGGGACAT	13020
	AAGCTCAAAT	TCCAGCACCA	GGAGATGTGA	CGTGAGAGTC	ACTTTTAGGG	GCAAGAACTT	13080
30	GATTCTTCAT	TGAAAGAGAA	CGCATTCCAT	GTGGATTAAG	TGCAGTTCTT	TCTGTAGCCA	13140
	GGGGAAAGAA	TGAGTTGAGT	TTTTGGGATC	CTCTCTGTCT	TTATGATTTT	ATGATTTTTT	13200
	TCCCCTGTTT	GATGCCCTGT	TCCCCAGACA	TATAGACCCA	GAATGACTCA	GTTCTGTTAA	13260
35	AGTAGGTTCA	ATCCAAAGTG	GGGGCAAGAG	ATGGGAGCGA	AGATGAGATA	GGAATCCAGG	13320
	AAGGCAGCAG	ATTCCAGAGG	CTTTCAAGGG	GGGTGGTGGG	TGGGTGTGAA	TGGGAACAGA	13380
40	GGGGATGGAG	CCAGTGGATT	ACAGAGGAGA	GAGGGAGAGG	AAAGAGAGAG	AGAGAGAGGA	13440
. •	ATGAGGGAGA	GGAGAGAGAG	GGGCAGAAAG	GCAGCTGCAT	GGATCTGGTA	GTTGGTACTA	
	AGAGAGAGAA	GCCGACAGAC	AAGGAGAGGT	TGAGGGGGAA	GAGGGAGATT	TGGGGAGGTA	13560

	GAGAGGAAAT	ACAGGCTCTA	CATCTGAAGA	AGGCAGTCTG	CTCCCTCCCT	ТТТАТТСТАТ	13620
	TCTTTGGGTC	TTCTATCCAC	TGTGTTCAGT	GGCCCTTTAA	TCCTCCCCCA	CTTTCACTCT	13680
5	•	ATTCTTCTCT	GATCCTTTGT	CTGTCTGCCC	ATTTGCCTCT	TGAGGTTGAC	13740
	ATCATGCTGT	CTGTCCCAGT	CCTTGCCTTG	TCTTTTCCTG	GTTCCTTTAT	GTTTCTTTAC	13800
10	CCCATCTTTG	CCTTCAGTGG	TAGGAGTGGG	TGAATGGAGT	GGCTTCCCCC	ACACAGAGCC	13860
	TCAGCAGGGG	CTCACCATTC	ACCTTCCCAC	TTGGAATCCA	CATCCTAAGA	CCAGATGCCT	13920
15	TCCCGAACTC	CTCACTTCAG	GGACAGAAGC	TGTTGAAGGA	AGGTTCAGAA	TGGCTGCTTC	13980
13	TTTGCTCTAT	CTGAGTATTG	CTCTGAAATC	CCCAGTTAAC	CTCTCTGGTC	TTTATTCCCT	14040
	CATGCACCCC	GTGTTTTTCC	AACTTGTTTT	TTATTCCCAC	CCAAGACTCT	GGACGACCTG	14100
20	GAGGAACGAG	TGAAGGAGGC	TGGAATTGAG	ATTACTTTCC	GCCAGAGTTT	CTTCTCAGAT	14160
	CCAGCTGTGC	CCGTCAAAAA	CCTGAAGGTC	AGATGGCTGG	GAGTGGTGGG	CTCTGTTTAC	14220
25	GGAGGGACCA	AGCTGGGGGA	CAGTGACTGG	TTGGAGAGGA	AAGCCAGGCG	GGGGCAGGTT	14280
_	TTGATTCTCT	GAGGCAATAG	CATCTCCTGG	GGAAGTTTAG	CTCCATCTTC	CAGTTGACGT	14340
	TTATTCACTA	TACGTTGAGC	GTTACCCTGC	ACTAAGCACT	TTGGGATGGG	AAATCAAAGC	14400
30	TGTGAAGACA	TCTGGCTTAG	CCCCTCAGGC	ATTCCCGGGC	ATCCCTCAGG	AGCTGTTTCT	14460
	TTCTCTGTTG	TAGCGCCAGG	ATGCCCGAAT	CATCGTGGGA	CTTTTCTATG	AGACTGAAGC	14520
35	CCGGAAAGTT	TTTTGTGAGG	TGGAGTTGGA	TCTGAAGAGG	GAGGGCACT	GGGTGGGAGT	14580
	TTCCCTTGGT	TTTCTTGTGG	GGCCTCCTCT	TGGCATCTGT	GCCTGAGTTG	ATAGCATATG	14640
	ATCTGAGGTG	ACGATTCATA	GGATGTCTCT	GTCTGTTGGC	TCTGACTGCA	TCCCTTGTCT	14700
40	GCACACACAT	GATACTTTCT	TCAGATCTCA	TTTTTCTACT	GCTTTGTGTT	TCCTGAGAAG	14760
	CCCATGAATT	CCATCTGTCC	TGACTGGCTG	GAAAAGGCCA	CTCAGAAATA	CAGGGGCTGG	14820

CTCTTTTTAG TTCTACTC	CT CTTCCCTGAT	TCCCAAGAGG	CTAAATAGTA	TCAAGTGAGA	14940
5 TAAGACAAAA ACAAACAA	AAT GAGCAAACAA	AAACTCAGCC	ATTCTCCTCT	GTATTCAGGT	15000
GTACAAGGAG CGTCTCT	rtg ggaagaagta	CGTCTGGTTC	CTCATTGGGT	GGTATGCTGA	15060
CAATTGGTTC AAGATCTA	ACG ACCCTTCTAT	CAACTGCACA	GTGGATGAGA	TGACTGAGGC	15120
GGTGGAGGGC CACATCAC	CAA CTGAGATTGT	CATGCTGAAT	CCTGCCAATA	CCCGCAGCAT	15180
TTCCAACATG GTGAGAG	TGT GGGGACTTGC	AGTCTGGCAC	CTGGGAGGGT	GGAGAGGACT	15240
15 GAGGGGGCCT TGCAGGGG	GAA AGGGTGGCAG	GGAGAGGGTG	CGGAATTTGG	ATATAAAGGA	15300
GAAGAGGGG CTGTGCCC	CAC CCTGAACTTG	TCTGCATTAT	GTTTCCTGTG	GATCCTACCT	15360
TTGCTCTGAC TTCCTTGC	GGT AGAGAGAGAA	ааааааааа	ACGATGGAGT	TGTATGTTCA	15420
GTAGGTTCCT GATGAGTC	GGA AGGGCTGTTA	CCATGGAGAC	GAGGAGCAGT	TGGTGAGAAG	15480
TCAGGAGGAA CCGGCAT	TAA TGATAATATG	GATGCTTGTA	TACTCAAGCA	CACCTTTACA	15540
25 GGAGCACTGT GTCTGGGG	CAG AATTGCATTT	CATTTTCTTG	GTGATTTATG	TTAGTGTTTT	15600
AGAGTTGCTT AATATTC	ACT CATGATTGAT	ATGCAATTAG	CTTGGATCCT	GTTGCTTTTT	15660
TTAATGATTC TTTTTTT	AGA ATTTTATGTG	GAGAAGGGGC	TTTTGAAATC	ATTTAGCCCC	15720
AGACAGCTGG TTAATGA	CAG ACTTGGAGCG	AGGAGGCGCA	TTCTCTGATT	TGAGAGGGCT	15780
GGGGCAGCCC TGCAGCT	GAG GGGCAGAAGT	GAGTGTGGGT	GTGGGCACAG	GGGAAGAGGC	15840
35 TTCTTTGTTG CTCAGCA	CTA CATTGTAAAG	AGTACAGGAA	TTTTGGGTAA	GCTCGATACG	15900
GCCACCTATG TGTCTGC	TTA TTACACCAAG	TGTGTACCTG	CTTTGGAGAG	CATGTTAGGG	15960
GTGAGTTAAT GGTTGTA	AAG CCCCAAAATA	GCTTGATTTA	AAAAGTTTTA	CTAACAGTGG	16020
CATTTGTGTG GTACTTA	GAT TCATGATTTC	AGGGATTCTA	ACAGCAAAAC	TCTGATGTGG	16080
ACTTATCTAC ATTTTGT	AGA CCAGGAAAAT	GAGACTCAGA	GTGGTTTAAT	GACTTGTCTC	16140

AGGTCATGCA GCTACTCAGC AGTCCAGCTG GAATTGAATC CTAGTTAAAT CTGGCTCCAA 16200 ACTGCTCTGC TCATCACTTC CCATGGAAAA CCATGAAGCC TGACTAAAAA GTTTGTACTT 16260 5 TTATCTGCTA ACACTGTTTA GCTACTGGCA ATTGCTGAGC AAAGGAGAAA TAAGATTTAG 16320 GAAGTTGACT CTGGTGGTGT GTTGTGATGA AAGATCTGCT GCCACATGGA CTCTGATGAT 16380 ACACACTGTT TTGGGACATT CTAAATGGAC TCTATGTTTT AAGGTATGTT AGTTGAAAAA 16440 TACAATGAAA AAATTTTGGA AGATCACTTT CCTGTAGAGT ACCCAAGAGG TGAGTCATCC 16500 ATAAAACCTG GGAAAGTATT GGAACTGGCT ACCAAAGCCA GAAAGACTTG TTGGTGCCCT 16560 15 GAGCCCAGGA CTTGCCAATC GTTGCTTCTG TCTTATAGCA GGGAGCTAGA GGAGGCTGAC 16620 AGAAAACAGG GATTTGGCTG GATCTAGTGG CTCACACCTG TAGTCCCCAG CACCTTGGGA 16680 GGCCAAGGCA GAGGATCACT GGAGCCCAGG AGTTCAAGAC CAGCCTGGGC AATATAGTGA 16740 GACCCCCAGC TCTAAACAAA AAGAGAGAGA GAGAGAACAG AGAGATGGTT AGGGACATAG 16800 AGGGACACCT GAGCATTGGG GTGGGAAGAC TTGGAAGGTC CCTTTCTGAC TTTGTGTCTA 16860 25 16862 GA

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- (2) INFORMATION FOR SEQ ID NO: 61:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 16707 base pairs

(B) TYPE: nucleic acid

- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)

10

- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: NO

## (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

### (ix) FEATURE:

- (A) NAME/KEY: intron
- (B) LOCATION:1..2742

### (ix) FEATURE:

- (A) NAME/KEY: exon
- (B) LOCATION: 2743..2985

## (ix) FEATURE:

- (A) NAME/KEY: intron
- (B) LOCATION:2986..3613

### (ix) FEATURE:

- (A) NAME/KEY: exon
- (B) LOCATION: 3614..3677

### (ix) FEATURE:

- (A) NAME/KEY: intron
- (B) LOCATION: 3678..5226

# 5 (ix) FEATURE:

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- (A) NAME/KEY: exon
- (B) LOCATION: 5227..5304

### (ix) FEATURE:

- (A) NAME/KEY: intron
- (B) LOCATION:5305..6848

### (ix) FEATURE:

- (A) NAME/KEY: exon
- (B) LOCATION: 6849..6999

## (ix) FEATURE:

- (A) NAME/KEY: intron
- (B) LOCATION: 7000..7494

## (ix) FEATURE:

- (A) NAME/KEY: exon
- (B) LOCATION: 7495..7627

(B) LOCATION: 11370..11591

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION:11592..11735

(ix) FEATURE:

(A) NAME/KEY: intron

(B) LOCATION:11736..12512

10

5

(ix) FEATURE:

(A) NAME/KEY: exon

(B) LOCATION:12513..13999

15

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 61:

AGCTAATTTT GGAATTTTTT TTTTTGAGAC AGAGTTTCAT TCTTATTGCC CAGGCTTGAG 60 TGCAATGGCG CGATCTTGGC TCACAGCAAC CTCCGCCTCC CAGGTTCAAG CCATTCTCCT 120 GCCTCAGCCT CCGGAGTAGC TGGGATTACA GGCATGCGCC ACCACACCCG GCTAATTTTG 180 TATTTTAGT AGAGACAGGG TTTCTCCATG TTGCTCAGGC TGGTCTCGAA CTCCGGACCT 240 25 CAGGTGATCC ACCCGCCTCA GCCTCCCAAA GTGCTGGGAT TACAGGCGTG AGCCACCGTG 300 360 CCCAGCTAAT TTTTGTATTT TTCATAGAGA CAGGGTTTCA CCATGTTGGC CAGGCTGGTC TTGAACTCCT GACCTCATGA TCTGCCCGCC TTGACCTCCC AAAGTCCTGA GATTACAGAT 420 30 GTGAGCCTCC GTGCCCAGGC CAAGTCTGGC TAATTTTTAA AAAAATTTTG TAGAGTTGGG 480 TCCTCTCTGT TTTGCCCAGT CTTGTCTCAA ACTCCTGGGC TCAAGGAATC CTCCTGCGTT 540 35 GGCCTCTGAA AATGAAAATG TTGGGATTAC AGGCGTGAGC CCCCTGTGCC TGGCTGCCTT 600 TTTTTTTTT TTTTTTAAA GGCAGAGTCT CACTCCATCG CCCAGGCTGA AGTGCAGTGG 660 CGTGATCTCT GCTCACTGCA ACCTCTGCCT CTTGAGTTCA AGCGATTCTT CTACCTCAGC 720 780 TTCCCAAGTA GCTGGGATTA CGGGAGCCCA CCAACACACC CAGCTAATTT ŤTGTATTTTT

	AGTAGAGACG	GGTTTCACCA	GGTTGGCCAG	GCTGTCTGGA	ATTCCTGACC	TCAGGTGATC	840
	CACCCTCCTT	GGCCTCCCAA	AGAGCTGGGA	TTACAAGTGT	GAGCCACTGT	GCCCAGCCTG	900
5	ACTTGTTTTT	TATAATGCCT	TTTTTTTTT	TTTTTGAGAC	GGAGTCTTGC	TCTGTCGCCC	960
	AGGCTGGAGT	GTAGTGGCGT	CATCTCAGCT	CACTGTAACC	TCCACCTCCT	GGGTTGAAGT	1020
	GATTTTCTCA	CCTCAGCCCT	CAGCCTCCTG	AGTAGTTGGG	ACTGCAAGTG	CACACCACCA	1080
10	TGCCCAGCTA	ATTTTTTGTA	TTTTAGTAGA	GATGGGGTTT	CACCATGTTG	CCCAGCTGGT	1140
	CTTTAACTCC	TGAGCTCAGG	CAGTCTGCTT	ACCTTGGCCT	CCCAAAGTGC	TAGGATTAAA	1200
15	GGTGTGAGCC	ACTGTGCCTG	GCCTTTTTTT	TTTTTTTTT	TTTTTTGAGC	AGTTTTAGTT	1260
	TCCCAGCAGA	ATTGAGATGA	AGGTACAGAA	ACTTCCCATA	TGCTTCCCAC	ATGCATAGCC	1320
20	TTCTACATTA	TCGACATCCT	CCGCCAGAGT	GGTACATTTG	TTACAACTGA	TGAACCTACA	1380
20	TTGATACATC	ATAATCACCC	AAAGTCCATA	GTTTACATTA	GAGTTCACCC	TTGGTGTTAT	1440
	ATATTCTATG	GGTTTGGACA	AATGTATAAT	GAGACGTATC	ТАСТАТТААА	TACTTTACAG	1500
25	AGTATTTTCA	CTGGCCTAAT	CCAATGGACA	TTTATTGTTA	CTTCATTATG	GTTGGGCACA	. 1560
	GTGCTAGATG	CTGATGATTA	AGAGAGGGCA	TGGGATTTGG	TCTTGTCCTC	AAGGGTAGAA	1620
30	CCTAGGCCCA	TTGCATCTTC	AAAGCCCAGG	CTCCTTCAAA	GCCCAGTGTA	GTAGCAACTG	1680
30	CTGTACCTTG	CCTGTGCCCT	TTGCGTATCT	CACTCCTCTA	TCTCTCTAGA	AAGTTGGAGA	1740
	GAAAAGTGAG	CAAGGCATGA	GGAACAAAGT	ТАТТТАТТТА	TTCTTCATTC	ATCTATTTAT	1800
35	TCTTTCATTA	CCGTTTGTGT	TAAAACATTC	CAAACCCAAA	CAATTATTTG	TATGGTCCCC	1860
	TGTGTATTAC	TTGTGGTTTC	CCAAGAAGTA	GTTGCTAAGC	TTTTCCTTGT	ATGGTTTCTG	1920
	TGAGGTAAGG	AAGGAATGAT	GTGATTTTCT	CCAGTATGTA	GAATGCAGTT	CCAAGAGGTT	. 1980
40	AAGTAATTTA	CTTACAGTTA	TTTAGCCAAA	CAAGGTTACT	GCAAGGTATA	TGAAGTCAGG	2040
	TCTCTTGACC	CAGTTCATGA	GAGAGTTAAA	GGAACTATCA	TTCTTTTTAG	CTTTCATGGA	2100

	AAAAGAAGGT	TGAGTGTTGG	GAGGGGTGTG	GGTAGGATTG	ATAATGGACT	TCAAAAATGT	2160
e	GAAGGGTATT	TCTGTAGTTT	TCATTCTTCT	GAAAGCCTTC	TAAGAGGCAG	TGAACCAAAA	2220
5	GCACACAAGA	ATGGCAAGAA	GTTAGCATGC	TGAAGAAATA	TCCTCCTGGC	TGGCAAGCAG	2280
	AGTGAGAAGA	CTGCTATCAC	CTTTTCTAGA	ATCTTTTGGA	ATTGTAGGAG	CTGTTAGATC	2340
10	CTGGGTTAAC	TCTATGAAGA	AAGTCAGAAG	GATCAGAGAA	CATCAGTGTC	ACAGCTCTTC	2400
	ATTGGAATAT	CCATGTCTCC	TCCTTTACTC	TGCTCTACCT	TCCATCCTTT	GCCACTAATT	2460
15	ATCCAGAGTG	TTTGTCAAAA	TTCTCTGTTT	GCAGTTCTGA	GCTAGCAACT	GTACACACTA	2520
	ACACCATCAG	ACACAGCTAA	TACCTACTCT	AGTCTAGTAG	CTTCCGATCT	AAGGCAGACA	2580
	CATGGGTATA	GTTAAAGATT	TTGAATGTAC	ATGTGTCCAA	TCTGACAACA	GTAACACAAA	2640
20	CCATCCATTC	AAGTAGAAGT	GATTGAGTCA	GAATTGGATT	GCACCCCTTC	CCCCACACCC	2700
	ACACACATTT	CAGTTCTTTC	CTCATGATTT	TTTCCTCCCA	AGACATCCCA	GGAATTTGTG	2760
25	GAGAAACTAA	CCAAGCGACT	GAAAAGACAC	CCTGAGGAGA	CAGGAGGCTT	CCAGGAGGCA	2820
-	CCGCTGGCCT	ATGATGCCAT	CTGGGCCTTG	GCACTGGCCC	TGAACAAGAC	ATCTGGAGGA	2880
	GGCGGCCGTT	CTGGTGTGCG	CCTGGAGGAC	TTCAACTACA	ACAACCAGAC	CATTACCGAC	2940
30	CAAATCTACC	GGGCAATGAA	CTCTTCGTCC	TTTGAGGGTG	TCTCTGTGAG	TTAAAACTTC	3000
	CTTCATACTC	CCCTGTCTTC	CCAATCTTGA	GAGAGACTCC	CAAGAGGCAC	CTTCTACAAA	3060
35	CATGCATTCT	CTGTTTTTCT	CAGTTACTTC	TTTGCAGAAT	CAGTCTCCGA	CCAGAGAAGT	3120
	AGGGACCTTC	AAATTAGAAG	AACCCATCAA	AGACTAGAGG	AAAAAAAATG	ATGTATTCCA	3180
	TTTTTTTAAA	CCCCTCCCCT	CATTTCTTTT	CAAACTAGAC	CAAGTATTCA	TGAGTCAGAT	3240
40	GAGAACTATA	GGATTTTGAA	AGACAAAACA	GTCTGAAAGG	TCATCTTCTT	ATTCCTTTTA	3300
	AAATGAAAAG	ATTAGTTTCC	AGAGAGATTT	GCTGACTTGC	TTAGGCCACA	CAACCAGAAG	3360

	CCTGCTGGTG	TTCTGTCTGG	GGATTTTTTC	CCATTCAAAT	CTCATAAGTG	AAGCTCCTTC	3420
	TCCAAAGAAT	AATGTTTCTA	AAATCTAGGG	TATGGGCATC	TGGGGTATGT	CCTATATGCA	3480
5	GGCAAATGCC	ATAAATAGCA	TTCATTCAGA	GGCTCAATTA	CATCAAAAAC	AGAAGGATTT	3540
	AAAGAGTCCC	TGATGTTCTC	TTTCACTCTT	GCTTTTGTCT	CCTTTGCCTT	GCTCCACATG	3600
10	TTCCTTCCCT	CAGGGCCATG	TGGTGTTTGA	TGCCAGCGGC	TCTCGGATGG	CATGGACGCT	3660
	TATCGAGCAG	CTTCAGGGTT	AGTACAGGGG	CAGGAGGGGA	CCGGACATGG	GGGCTAGGCT	. 3720
	GGGGCTGGGC	TGGGATGCCC	CCTGGGGAAG	AATGCCAGAG	ACATCACAAG	ATTGCCCTGG	3780
15	CACCTCCCAA	CTTCTGCCCT	TCTCTTTTAA	CTCTGTTCAT	CAAGCTTGTA	ААТААТААТА	3840
	ATAATAAGCT	TAACTACAAG	AAGATTGATG	TCTTTGAGTT	GCACTGGTTT	TGCTCTTGAA	3900
20	AAGAGGTGTG	CAGGCTGGGT	GTGGTGGCTC	ACCCCTGTAA	TCCCAGCACT	TTTGGGAGGC	3960
	CAAGGCAGGC	AGATCATGAT	CATGGTCAGG	AGTTTGAGAC	CAGCCAGACC	AACATGGTGA	4020
	AACCTGTCTC	TACCAAAAAT	АСААААААА	ТАААААААА	TAGCTGGGTG	TGGTGGCAGG	<b>.4080</b>
25	TGCCTGTAGT	CCCAGCTACT	TGGGAGGCTG	AGGCAGGAGA	ATCACTTGAA	CCCAGGAGGC	4140
	AGAGGTTGCA	GTGAGCTGAG	ATCACGCCAC	TGCACTCCAG	CCTGGGTGAT	AGAGTAAGAC	4200
30	TCTGTCTCAA	AGAAAAAGA	AAAGAAAAGA	GACATGCAAA	TTAAAAACAG	CTACTCTCTT	4260
	TCCCAGTGGC	TTCCATTAAT	TTCAGGAATT	TCCCCTTGAG	TGGCTTGGGT	TGAGAGGTTG	4320
	ATGACCTGTC	AGTTAGACTC	AAGAAAGCTG	AATCTAGGAG	AACCGCTATT	TTTTTTTAA	4380
35	GGGAATCTGC	CAAATTTCCT	TGCTGTGTAA	AGCTTCAATG	TGTATAGCTT	GGCTTTTGTA	4440
	GATTGTATTT	TCTTGAAACT	TAGCACACAG	GTATTTGCAG	AACTTCTAGG	AGTTAATTTT	4500
40	TCTGCTCCAC	TCGGCTCTCA	GTCTTTTACG	GCATGGCCAA	GAGAGCTATT	TCTTGGCCTC	4560
	CTGTGAAAAG	TTTCTTTCTT	CCTTTCTCCC	CACCTCCACA	TCCTTTCAGC	TCCTCTTTGT	4620
	ATCCAGGACA	AGAGGAAATG	GACTTCAGCC	ATGGTGAAAG	GAGTGTGAGT	TGGCTTTTGA	4680

	AGGAAAAGTT	ATGGTAACGG	AAACAGTTCT	AGAACAGAAA	TCTTAGAAAT	GACCAAATTT	4740
5	TACTCAATGG	CGCTTTAAGA	GGCAGATATA	ACTTATCCAA	GGAATTAAAA	CCCAAGCCAA	4800
,	CAGAAGAGAA	TGTTCTAAAA	ттаааатсаа	AGCCACTGGG	AAAATAGAGC	CTGCCCATCA	4860
	TGAGAGGAAG	AATAAGCAGA	AATATGTGTA	AAGCTTTAGA	AGCCAAAATC	AAAGTGAGAG	4920
10	ACATCTCGCC	GAGAGAGGTG	TGAGGAATGG	AATAGGTGGC	AGACATGTTG	TGGAGCCTCC	4980
	TCACTGAAGA	СТТТТАААСА	TAGATATTCT	TATTTATTTG	AGTTGTCTTG	GGAACCACCT	5040
15	TATATTGCTT	TTAAGTCATG	TTGCTGATTC	AAGAGTCTCG	TAGGTCCTTC	CAAGCATCCT	5100
13	TAGGGCCTCA	GGTGAAAATA	AAATCAGATA	CAACCATGCA	AAGCTCTAGG	GAAGTGGGAA	5160
	GTTGAAAATG	CCTAGGATCA	GCTCTTTGGC	TACCTGTGGT	CACTCCTTTT	ATTGTCGTCT	5220
20	GCCCAGGTGG	CAGCTACAAG	AAGATTGGCT	ACTATGACAG	CACCAAGGAT	GATCTTTCCT	5280
	GGTCCAAAAC	AGATAAATGG	ATTGGTGAGT	GGATCTTGTT	TGTATTTTCC	TTCAGCCCCT	5340
25	CTCGACAGTC	AAGGGAAAAA	GTCATGCCTT	TGAGTGAGGA	TGGAATGGTA	GAGACTGTTA .	5400
	GGTTGGAATG	TGGCTGGCAG	CTGGGCCAGG	AGAAAGGGTT	AAGTGAGAGT	GAATACAACC	5460
	CCTAAGGCGT	GGGTAGGGGA	GACTGGTGTA	TTTGGAGAGG	GAATAGGCGG	TGGTTAGTAC	5520
30	TATTTTTAAT	GGTGCATTGC	TGGGGTAACT	GGGGATTAGA	GGCAGGGGGT	GGGCAGAGGG	5580
	CGGGAAATGG	AAACTCCATT	TGGGTTTCCC	AGATGTCCTG	GTGTCTTGAT	ATATTTGAAC	5640
35	CAGCTACTTC	AAGCCCAGAG	CTGTCTCTTT	GTCTGTCTCT	GTCAGGAAAA	CGGTTGCTTA	5700
	AACTATGGAG	GAGGAGGGAA	AACCTCATGT	AATTGTCATC	TGCCAAAATG	TGCTTTTTAT	5760
	TTTTATATGT	ATTTTTAAAA	ATTTTCCTAT	TTTTATGTAA	TTTAGAGGTA	GACGTGCAGT	5820
40	TGTGTTACAT	GAATATATTG	CATAGTGGTG	AAGTCCGGGC	GTTTAGTGTG	CCTGTCACCC	5880
	GAACAGTGCA	CCTTGTACCT	AATAGGTAGT	ATTACATCCC	тсаааатата	CŤTTTTAAAG	5940

	AGAGAAAGCA	AGCAGTTATT	CTTTGTGTAC	TTGGTCTAAA	TGATAGGACA	TAGGAGAGAA	6000
	ACTGAAGGTG	GACAAAAGGA	AGGACCTACT	GATAAAAGAA	AGCCTCCTTG	AGAATGAAGG	6060
5	GGAGGCTCAA	CCATTGAAGA	TGGCTGCCGT	CTGCCCTGCC	CAGCAGATAT	CCAGTCATTC	6120
	CCAGCACTGC	TGGAGTTTTG	CCCTTTTTT	TTTTTTTTA	CAATTCGAAT	TTAGGACAAT	6180
10	GTTCTGGATT	GCTATAAATG	CTGCATGGCC	TAAATTATTC	ТТТААААААА	AAACTAAGCA	6240
10	AATTGAAATT	AGTTTTTTT	GGTGAACTCT	GACAAATTTG	AACTTCCCCC	ТААТААТААС	. 6300
	TGGAAAACAT	ATTTGGGAAT	ATTACCCTGC	CAGGATTAAA	ATTTCAGATT	AGCTTTCCTT	6360
15	CTTTTTTGT	TTGTCTTAAG	AATAGGTGTC	CACACTAGAT	ACTTCAAGGC	CTTTTTAGCT	6420
	TTATGATTCC	ATAATTGTCA	TTTAAAACTT	TGATTTGGGT	TATAAGAAAC	CTTATAACAT	6480
20	TTTTTAATGA	TCCCCTTCTT	TCTCCTCCCA	TTTTCCTTTG	CTGTAAGAAA	GACAGAAAAA	6540
20	CTTAAAGAAC	AAACAAAAAC	AAAGACTACA	ACTTTGGGGA	CATGCCTCAG	CATTTCCCAA	6600
	CCTATGGATA	GACCATTCAC	TCCATCTTCT	CATCTCATTT	CTGGTTGCTT	CCTAACGGCC	6660
25	CCAGTGGCAC	TGAGCATTCT	GCCTGCAGTA	ACCTCTGTCC	AGTGCAGTTA	GGGCCTCATG	6720
	TCCCCAGCCA	ATGACTGAAT	GTCCATCAGC	AATCTAGTTC	TTTGCCCTTT	TCTCCTATCC	6780
30	CGTCTTCATT	CCTTTGTCCT	CCTTCCCTTC	TCTTTTCCCT	TCCCCTCTTC	CTCCCCTGTG	6840
	CCATGCAGGA	GGGTCCCCCC	CAGCTGACCA	GACCCTGGTC	ATCAAGACAT	TCCGCTTCCT	6900
	GTCACAGAAA	CTCTTTATCT	CCGTCTCAGT	TCTCTCCAGC	CTGGGCATTG	TCCTAGCTGT	6960
35	TGTCTGTCTG	TCCTTTAACA	TCTACAACTC	ACATGTCCGG	TAAGTTTCTC	TTCTGACGTT	7020
	TTCCTTGTCT	GCCTCTCTGA	GATACTGATC	ATGTTCCTGG	ACAGGATGAG	ААТААААССТ	7080
40	GTGTAACTCC	CATGGCCATG	TATCATGGAG	TTTTTCATTC	TGACTTGTTG	AGAATGAAAA	
	CAGGGAAACC	AGATATAACC	CCCACTCCTA	CTCCAAAGTA	GCTAACGGGA	GGAAAAAAGA	7200
	AAAGAAGAGA	AAAAAACAAC	CTTTGGGGCC	AGGTCTCACA	GTCTTGGACT	СТАСАТАААТ	7260

	AGCCTGTATT	CTAGTGGGGG	CCTGTGCTTG	GGAAGCCCTC	TGCAACTCCA	TCTTCAGCCC	7320
5	CATGACTGCA	TTGCTCTGCC	TCTCAAGGCT	CCACTGTCTT	CTCCAATCCT	GTCTTCCTTT	7380
•	AGCCCCTGGC	CCTGAAATTA	GGGTCATGCC	ATTGCGTGGT	ATTTGGAGAG	CTCAGCCTCC	7440
	CTGGAGAAGA	GGGGTAATTC	TCTCTCCCTC	TCACCCTCTC	CACCTCTGCC	CTAGTTATAT	7500
10	CCAGAACTCA	CAGCCCAACC	TGAACAACCT	GACTGCTGTG	GGCTGCTCAC	TGGCTTTAGC	7560
	TGCTGTCTTC	CCCCTGGGGC	TCGATGGTTA	CCACATTGGG	AGGAACCAGT	TTCCTTTCGT	7620
	CTGCCAGGTG	AGGAGGTGGT	GGGCAAATTC	CTTACAGGAT	GTGACTCTCC	CACCCGTCTC	7680
15	AGGAGCACCT	TCCATGATTT	ATGATTCTCT	GCCCTTCCTC	CTCAGCTTTC	CCTGACTCTT	7740
	GTCCCTGTTC	TTTCCTTCTA	GCATCACCCC	TCTGTTCTCT	GTTTGGCTCT	GTCCCTTCTT	7800
20	TCTGTGTCTG	CAGGCCATTT	TCATTCTGTA	GTTTACTTGT	CAGTTCCAAG	GTTGCCATGG	7860
	CAGSCCTYGC	AGAGAAGAGG	AGGGAGCCAT	TGAAGGCAAA	GGAAGGGGAT	CTGCTCAAAG	7920
	GTCTCCTGAA	CAATGGTGGC	TTGTCTGTGG	TATGGGGGCT	GAGAATCAGA	ACTGTGGACT	7980
25	TTTTTTGGGA	GCCTTTGTTG	GGTTTGGAAG	GATAGAAGCA	GAGATGGAAA	CACAGCAGAG	8040
	AGTTGGGGGG	AAGGGACCAC	TGCCACACAG	GGGAGGAGGG	GCTCTGGGAC	TGTTGGTACA	8100
30	TGGAAGGTTC	TAGTGCTGTG	GGGAGAGGCC	AGCTTCAACA	GTGATAGTTG	AGTGGTTCTC	8160
	TTTTCCACTG	GTGGAAACAC	CCACTCTTTC	TCCTGATCTG	CCTGCCTGTC	CTTGCTCTCT	8220
	CTTTTTCCTC	TGCTCTGTGC	TGTCCTGATC	ATACATCTGT	GCACATGGCA	TTTCCATGCA	8280
35	CATGCACATG	CAGTTCATCA	GGAATCCTCT	GTTCCCAGTG	AGGCCAGAGT	GCAGCTGGAG	8340
	AAGCAGACAA	TTAGCTGTAG	TGCAATAGGA	GAGGTTCCAG	AGTAGGGATC	TGCACAAAGT	8400
40	GCTTTGGGGG	CAAAGAAGGG	AACACAGTTC	ACTGCTGGCG	TGATTGGGTG	GACCTCACTG	8 <b>4</b> 60 <sub>v</sub>
	AAGAGGTGGC	ATTTGAATAC	TGAAGGACAA	ATAGGATTTT	ATCAGCTAGA	<b>бааатадад</b> д	8520

	AAGGCTACTT	CAGGGGCATA	GGGAGCATCG	TGTGGCTAGA	AAATACATGA	AAGAGAGTAG	8580
	ATGAAGAGAA	AGTGAGTAGT	TCAGCATGGC	TGGAGCGTGG	GGTAGGTGTG	GGGCTGGGAG	8640
5	ATGAGCCTAG	CTGGACAGGT	GGATGGGAGC	ATGTTGTGAA	GGGTCTGTGT	CATATCCAGA	8700
	AGTGTTCAGG	СТАТААСТТА	TAGATATTGG	GGAGTGGTTG	GAGGTTTTTG	GCCACTAAAG	8760
10	CCAGGAGGTT	TTAGCAAGAT	CACCCTGGTG	GTGTGGAAGT	AGAGGGTGGA	TGGGAGGAAT	8820
10	TGTTCAAGGT	GGGGAGACTG	СТСТССТССТ	GCCGCTCCCC	GTCCTGCTCA	CATTTTCGCA	. 8880
	TCCTCCCTGT	GCCACCATGA	GCTCCCTGCC	CGTGCTCCCT	GCCCACTCTC	CCTTAGGGTT	8940
15	CTGCCCATCC	TTACTGCAGT	CCCGGCTACT	ACTCTACCCT	GTTCTGCCTG	TGCCCTCTCT	9000
	TCCTTTCTAG	GCCCGCCTCT	GGCTCCTGGG	CCTGGGCTTT	AGTCTGGGCT	ACGGTTCCAT	9060
20	GTTCACCAAG	ATTTGGTGGG	TCCACACGGT	CTTCACAAAG	AAGGAAGAAA	AGAAGGAGTG	9120
20	GAGGAAGGTG	AGCTGCTGCC	CAATCCTCAG	CCCCCARATC	CTTGGCTCCT	GGGGCACAGA	9180
	GCATTTTCCC	CTGACGTGCC	TGTTCTCCCC	ACATATTTAT	CCAGACTCTG	GAACCCTGGA	9240
25	AGCTGTATGC	CACAGTGGGC	CTGCTGGTGG	GCATGGATGT	CCTCACTCTC	GCCATCTGGC	9300
	AGATCGTGGA	CCCTCTGCAC	CGGACCATTG	AGGTACCACT	GGAGAGGAGG	TGCTATGGTC	9360
30	AGGAGAATGA	GCAGGGCTCA	GTGGCCATCA	GGGCCCTGGG	GCTGTGTGTG	TCTTGAGGGA	9420
30	TGAAGCTACT	TGGAGAGAGT	GCCTTCCTCG	TATTGGAAGC	TCTTCCTTTC	CTTCCTAGAA	9480
	GGAGCCCCTC	ATAGGCCTCC	AGATTCAGCT	GAAGAAAGGA	AGGGGTGGGA	ATCTGGGAAG	9540
35	GGTGTGTAGA	ACTTCCAGGC	ATCAGGGAAA	GTGGGGAACA	AGCACCTCCA	AGGGTTCAGG	9600
	AAAACATTCT	TAGGCCTAGA	ATGAGATTTG	GCATCAGCAT	TGAGGGTCTC	ATAGGAAAAC	9660
40	AGTTGGAAGC	CAGAGACTGA	GAAGCGTTGA	GGAGAGGAGG	GGAGGCTGGC	AACCATCTTT	9720
<del></del>	CTTGTGACCT	TGTTTCTGCC	CTAGACATTT	GCCAAGGAGG	AACCTAAGGA	AGATATTGAC	9780
	GTCTCTATTC	TGCCCCAGCT	GGAGCATTGC	AGCTCCAGGA	AGATGAATAC	ATGGCTTGGT	9840

	GTGTGGGATG	TGGGCAAAGG	AGGGCAGGGA	TGCACAAAGG	CAGGAGGGAA	GGCAGGGGTA	9900
5	GAGGGCTTGG	AGGGAGAGGG	GTCTTTGGAA	GAGGAGGTAG	AGAGCTTGTC	AACCCAGTTT	9960
3	GAACACCCTA	CTCTTTGTTA	TTGCACTAAT	CTTTTCTGAG	AATAGGGGAG	AGTTGCTCTT	10020
	TTGCTATGAG	GAGCTTAGGG	CCCAAAGCAC	AGAAAGCACA	GATGAAGAAC	TTGTGTTCAG	10080
10	CAGAGGAACA	AGTGGGGGTA	ACCCCACCTC	CAGACTTGAC	ATTATCTTTT	AGATCCCCCT	. 10140
	TGGCCTTATT	AGCATTGTTC	GATTCATGGT	CACAAATTGC	AAACCTACCC	TCTGCCTGGA	10200
15	AAGCCACCTT	CCCACCTGTA	GGGTAAGGGT	GAGGCATGTG	TGGCCCAGAC	TGGCCTATTT	10260
.5	CTAGATATTC	AACAAGCCCT	TGCCTGACTG	ACAGCAGCTT	GCCACCATTG	CTTTCCTGTG	10320
	TGAATCCCAG	GAAAAAGTGA	TGTGGTCTGG	GCAAGTTGGG	TGGACATAAG	GGATAGGGGA	10380
20	CACAGGGTGA	GGTTTGCTAG	GTCAGAGGGG	TTGGATTGGA	GAGGAGGCC	CCCTTTCCAT	10440
	TTCAGAGTAG	GTGAAGGGCA	GAGAGGGGAT	GGGGATTGAG	TGAGGAGCAT	TGTGGTCCTT	10500
25	GTTGCTCAAG	TGACTCTCTC	CTGCCATCCT	AGGCATTTTC	TATGGTTAAC	AAGGGGCTGC	10560
_	TGCTGCTGCT	GGGAATCTTC	CTTGCTTATG	AGACCAAGAG	TGTGTCCACT	GAGAAGATCA	10620
	ATGATCACCG	GGCTGTGGGC	ATGGCTATCT	ACAATGTGGC	AGTGAGCACT	GACCCCATGG	10680
30	CATTGACCCT	GTAGGCTGAC	CACAGCAGCC	CAGATATAGA	GGACTAGGAA	GAATCAATGC	10740
	TAGATCTGGG	ATCGGTTGCT	TAGAAGTCTT	AAAAAGTTTG	TTAATTCTTC	AGGTCTATAA	10800
35	AGCACTTTAC	AGTTTACAAA	GCTCACTACA	GACATTGTAT	CATTAATCTT	GCAACTACCC	10860
	AGTGAAGTAG	ATATTAGTAT	CCCCACTTTA	TAGGTGAGGA	AACAGAAACA	CAGAGACGTT	10920
	AAATTGCTTG	TCTGTGGTTA	ATGGGCTGGA	CTCTATTGAC	ATTTCCTGCC	AGGGACCGAC	10980
40	TCTGGAGGAC	CCGGAATCTG	TGCATAGAGA	TCCTGGGAGT	TCCTGCCTTG	AGGGGAGGG	11040,
	TTAACCAAGA	GTGAAAACTG	GTTTGGGACA	GTTTGAGATT	TTTCTCAATC	TATATTGAGG	11100

	ATGATCCTGA	ATTTGGATCC	TTTTCAAAGG	GAAAGTTCAC	CAGGAAACTG	TCTGCATAGA	11160
	CTCCCTCCCA	TGGGAAGTAA	ACTCTGGATC	TTGTCTGAGC	CTGCAGACCT	GAGACTCCCT	11220
5	CAATGTGTCT	TTCCCTCTAG	GTCCTGTGCC	TCATCACTGC	TCCTGTCACC	ATGATTCTGT	11280
	CCAGCCAGCA	GGATGCAGCC	TTTGCCTTTG	CCTCTCTTGC	CATAGTTTTC	TCCTCCTATA	11340
10	TCACTCTTGT	TGTGCTCTTT	GTGCCCAAGG	TAAGGATCTG	GCTTTTCTCC	CACCCTCTTT	11400
10	GTTCCCATGT	TCCCTCCATC	CCTCCTTCCT	ATATTACTGA	GTTCCTCTGC	CCTTCCGTTC	11460
	ACCCTCCTCT	CACTCCTCCC	CTTGTTTTGG	GCCCAACTCT	TATCAGCATT	CCTTCCACCT	11520
15	CCAACCTTCC	ATCAGCCAGT	CACTAGTACA	GTCCTTGCTG	GGCCACCCCA	CGCCCAAACA	11580
	TTTGCCCCCA	GATGCGCAGG	CTGATCACCC	GAGGGGAATG	GCAGTCGGAG	GCGCAGGACA	11640
20	CCATGAAGAC	AGGGTCATCG	ACCAACAACA	ACGAGGAGGA	GAAGTCCCGG	CTGTTGGAGA	11700
	AGGAGAACCG	TGAACTGGAA	AAGATCATTG	CTGAGGTGCG	GGGGTGGGTG	TCAGGGTAGG	11760
	GTGTTGGAGT	GGTCCAGGAG	GCTTGCGTCT	TAGCTTGGGT	TGTCTGAAGC	CAAGCCTGAG	. 11820
25	ATACAGGGTC	AGATGTTCTT	GGCTCATGGA	GGGAGGGTCC	TAGGAGACAA	CCTGTAAGGA	11880
	GTGAATGGAG	CAGCATAGGG	GAGGGGAAAG	GGCTGAGCAA	GATTCTATCT	CAGGCAAAAT	11940
30	CCAGTGTTGG	CCTGGCAGGT	GGAAGGGCTC	TGGAGTGGGA	GCTATGTGGT	TGACTCAGCC	12000
	TCCTTAAGGC	AAGAGGATGG	CTGTTGGCTG	TAGGTGACAA	CTGGAGAGAG	GCAGCTGTGA	12060
	GCCTCTAGTA	GTCAACACTC	ACAGCAGCTG	GGTGTAGCAT	GCAGCCCCAG	CATAAAGGAC	12120
35	CTGGGCAGGC	GTTCACTGTG	CCCCAGGCTG	TCATTAGGGG	CTGGTGCAAT	GCCAAAGAGA	12180
	GGGATGTTCC	AACTGGGTTG	ACACATCTCT	CTGATTTATT	GGAAGCTCTG	TGCACTGACT	12240
40	TTTCTCTCCT	TCCCCACTTT	TTCCTTTTGT	TTTTAAATTC	TCTCTTATTT	CCCTGATCGC	12300
	ATTTTTTCTA	TCGGTATCCT	TATGTTCTCT	GGCTTTTCTT	GTTCTGTTTT	GATTTCTCCT	12360
	TTTAATTTAT	TCTGTCCACT	TACCCTACGT	CCTCCCCTA	CATTTTTCTG	TGCCCTTCCT	12420

	CTCTTTCCCT	GTGCCCTTCC	TCTCTTTCCC	TCCTCCCCAC	TCCTTCATCA	CCTCCTCTTC	12480
	TCCTACTATC	CCAATTGTGC	TTCTTCCTCC	AGAAAGAGGA	GCGTGTCTCT	GAACTGCGCC	12540
5	ATCAACTCCA	GTCTCGGCAG	CAGCTCCGCT	CCCGGCGCCA	CCCACCGACA	CCCCCAGAAC	12600
	CCTCTGGGGG	CCTGCCCAGG	GGACCCCCTG	AGCCCCCGA	CCGGCTTAGC	TGTGATGGGA	12660
10	GTCGAGTGCA	TTTGCTTTAT	AAGTGAGGGT	AGGGTGAGGG	AGGACAGGCC	AGTAGGGGGA	12720
	GGGAAAGGGA	GAGGGGAAGG	GCAGGGGACT	CAGGAAGCAG	GGGGTCCCCA	TCCCCAGCTG	12780
15	GGAAGAACAT	GCTATCCAAT	CTCATCTCTT	GTAAATACAT	GTCCCCCTGT	GAGTTCTGGG	12840
13	CTGATTTGGG	TCTCTCATAC	CTCTGGGAAA	CAGACCTTTT	TCTCTCTTAC	TGCTTCATGT	12900
	AATTTTGTAT	CACCTCTTCA	CAATTTAGTT	CGTACCTGGC	TTGAAGCTGC	TCACTGCTCA	12960
20	CACGCTGCCT	CCTCAGCAGC	CTCACTGCAT	CTTTCTCTTC	CCATGCAACA	CCCTCTTCTA	13020
	GTTACCACGG	CAACCCCTGC	AGCTCCTCTG	CCTTTGTGCT	CTGTTCCTGT	CCAGCAGGGG .	13080
25	TCTCCCAACA	AGTGCTCTTT	CCACCCAAA	GGGGCCTCTC	CTTTTCTCCA	CTGTCATAAT	13140
۵	СТСТТТССАТ	CTTACTTGCC	СТТСТАТАСТ	TTCTCACATG	TGGCTCCCCC	TGAATTTTGC	13200
	TTCCTTTGGG	AGCTCATTCT	TTTCGCCAAG	GCTCACATGC	TCCTTGCCTC	TGCTCTGTGC	13260
30	ACTCACGCTC	AGCACACATG	CATCCTCCCC	TCTCCTGCGT	GTGCCCACTG	AACATGCTCA	13320
	TGTGTACACA	CGCTTTTCCC	GTATGCTTTC	TTCATGTTCA	GTCACATGTG	CTCTCGGGTG	13380
35	CCCTGCATTC	ACAGCTACGT	GTGCCCCTCT	CATGGTCATG	GGTCTGCCCT	TGAGCGTGTT	13440
33	TGGGTAGGCA	TGTGCAATTT	GTCTAGCATG	CTGAGTCATG	TCTTTCCTAT	TTGCACACGT	13500
	CCATGTTTAT	CCATGTACTT	TCCCTGTGTA	CCCTCCATGT	ACCTTGTGTA	CTTTCTTCCC	13560
40	TTAAATCATG	GTATTCTTCT	GACAGAGCCA	TATGTACCCT	ACCCTGCACA	TTGTTATGCA	13620 <sub>9</sub>
	CTTTTCCCCA	ATTCATGTTT	GGTGGGGCCA	TCCACACCCT	CTCCTTGTCA	CAGAATCTCC	13680

	ÄTTTCTGCTC	AGATTCCCCC	CATCTCCATT	GCATTCATGT	ACTACCCTCA	GTCTACACTC	13740
	ACAATCATCT	TCTCCCAAGA	CTGCTCCCTT	TTGTTTTGTG	TTTTTTTGAG	GGGAATTAAG	13800
5	GAAAAATAAG	TGGGGGCAGG	TTTGGAGAGC	TGCTTCCAGT	GGATAGTTGA	TGAGAATCCT	13860
	GACCAAAGGA	AGGCACCCTT	GACTGTTGGG	ATAGACAGAT	GGACCTATGG	GGTGGGAGGT	13920
10	GGTGTCCCTT	TCACACTGTG	GTGTCTCTTG	GGGAAGGATC	TCCCCGAATC	TCAATAAACC	· 13980
10	AGTGAACAGT	GTGACTCGGC	ACCTTGCAGT	CTTCCTGTGA	ACAGAATGGG	CTTCAATCCA	14040
	AGAAGGGAGG	CTCAGAGGAC	TCCAAGTTCA	TGAAAAGGCA	TTAAAGCGGA	GGGTGAAAAG	14100
- 15	AGGTGTTTTA	TTGATCCATT	GAGGGCTTAG	CAGAATGAAG	CAGGACATGA	TTAAGTCTGA	14160
	GATTAGTGAG	TGAGGACACT	ACTGGTTAAA	AGTGTGGGCT	CTGGAGTCAG	ACTGCCAGGG	14220
20	TATCAGATCC	AACCACATGC	AAACATTTTC	TTAGTCTCTA	TTCCCCATGT	CCTCATTTAT	14280
20	GAAAATGAGA	ATAACAGTAA	TACATTCCTC	CATAGGTTGG	GTACAAAGAC	ТАТТАТАААТ	14340
	TGTGCATTCA	GGTGCCTAGG	TTGGCCCTTG	GGCCATGGTA	TATGTTGCGT	GAATGTTAGC	14400
25	CTCTGTCCCT	GCTGTTTAAT	GAGTTCCTTG	ACAGTAGTGG	GCATGTATTG	GGAGCCTGGA	14460
	GCAAGTGCCT	AAGCATCCCC	TCTAGGGACG	CTCCTTCCCA	GGAACTAAGA	AGAGTAAAAG	14520
30	AATGATGACT	GCTAGAAGGT	AATGGATGAG	ATGGCTGCTG	AGTGCTTCCA	ACCTTAAACA	14580
	TCTTTGTTTA	GGAACTCTGA	GCATCTTGGA	AATAATTTGC	TATCAAACTG	AAAAAAAATC	14640
	TTGAATGGAC	AAGGGCAAAA	ACATTTGCCT	GAGACTTTAA	ACATTTTTTG	TGTCATCTTG	14700
35	GAGAGTGCTT	TTTTGAAGCT	CAAATTTTCT	TTTATTTTGG	CACTGATTTT	TAAAGTGATT	14760
	CTCAGATTTT	TGTAAGACAG	CTGCAAGGGT	TGGATGGGCC	CTGTCATTCA	CTGACCTGTT	14820
40	AAGAGCCAAT	TTCTGAACTT	CCACTAAAGC	ATGGGCTGGT	TGAATCTTAG	ACCAGTACTT	14880
70	GAAAAACTTT	CCACTGTGAT	TATCCACCTT	GGACCAGTTG	GACTTAATTT	GAGTCTCTCT	
	TCTTCCCACA	GTGAAATATC	CGCTAGGAAA	AGAGAAGAAG	TCTATGGAGT	AGATAAGCCT	15000

	GCAGTTTGGA	AAATGAATAG	TTGGCATCAG	AACAGCAGCA	GGAGGCTGGG	TGTGGTGGCT	15060
_	CACGCCTGTA	ATCCCAGCAC	TTTGGGAGGC	TGAGGTGGGT	GGATCACGAG	GTCAGGAGTT	15120
5	CAAGACCAGC	CTGGCCAAGA	TGGCGAAACC	CTGTCTCTAC	ТАААААТАСА	AAAATTAGCT	15180
	GGGCGTGATG	GTGGGTGCCT	GTAATCTCAG	CTACTCGGGA	GGCTGAGGCA	GATAATTGCT	15240
10	TGAACCCAGG	AGATGAAGGT	TGCAGTGAGT	TTGTGCCACT	GCACTCCAGC	CTGGGCGACA	15300
	GAGTGAGACT	GTCTCAAAAA	ААААААААА	GAAAAAAAA	AAAGAAACAA	CAGTAGCAGG	15360
	AGCTATAGAA	CAGCCCTGGG	TAGAACCTAA	AAGACCCAAA	TTATCATCTC	AAACTTGCAT	15420
15	TGCACTTAAG	TGGGCTGTAA	ATTATAAACA	AAGGGTGAAA	AGTTCTACTG	TGGCCAAAGG	15480
	TAAGCCAGAC	ACTCTGCTAG	CAGGAGTGCA	GGAGTCGAGA	GCCAAACGGT	GCGGCTAGCA	15540
20	GAGTGCCCAG	TGCAAAGGGG	TGGGAAGGAG	TGAGATTGAG	AATATTAAAA	AGGTACTTAG	15600
	AAGAGAACTT	GTAAGATTTT	TACTGGCCAA	ATTTAAAACA	TGACTGAGCA	CTATTTTCA	15660
26	TACAGGCCTC	СТАСТААТАА	GAAAACAATT	TTGAGATAAC	TACTTATTTG	AGTTCACAGT	15720
- 25	TAATGTTCCT	GATGATTAAG	ATCAGTTGCA	AATGTTCATC	TGTCAATGCT	TATCTACAAT	15780
	GAGACTTCAT	GTATTCATTT	CTGAAAGTGT	CTTTTCAGGG	TGAATGGTGC	TATTGATTAG	15840
30	САСТААТАСТ	AATTATTAGT	ACATTATATA	ТААТТАСААТ	GAGATATACA	CACACATACA	15900
	CACACACATA	ТАТАСАТАТА	CACACACACA	ATGATATATA	АТТААТАТАТ	AATTGTACCC	15960
25	CAAGGGGTGC	AAAGGAGATG	TGTTGCCAGG	TGGAGAGGCT	ACCGCTTGGC	AGTTCTGGGA	16020
35	GGACTTGCTC	CCTGTGCACT	GTGAGGCAGG	CTTTGCCTTT	CAGACCTGCC	TTTGGGTAGG	16080
	GTTCAGATCA	CTTTCTAACT	CTGGAATGTC	CTAGAATGTA	GACTGCCTGC	AGGCTTCCAT	16140
40	GTCCCCTGCT	TTTCCCTGAC	TTAGCCTGTT	GCCTCCCTGC	CCTCCTGTTG	GTTGTCTACC	16200,
	AGTAGAGAGC	ACTTTGTGTG	CACTTGGCTG	CTACATTAGT	TAGGTGATCT	TCAACAAGTG	16260

TTGATGGTGA GTTGCTGTGG CAGGTGCTTT TTTTGGCACT GAGGCAAAAT GGTGAGTAAG 16320

ATGGCTTTCA AGCGTTGTAC CTTCTCGACG TGGGAAGATG ACCAGTAAGC AGAAAAACAA 16380

5 ACGAGATCAC TTAAGAGAGC AACCAGGAGT GTTGGGCATC TCACAGCCAT TAGCTCTGGT 16440

GTGAAGGACA AATCTAAAAG CAAGGGGACT GTGTGTTCAT TTTCTGGGGT CACAAAACTA 16500

AGGAGCAAAG CCAGTATTCA AACTGCATAT ATTATTATCT ATTGCCACAA AACCTGTTAC 16560

CCTAAATGGC TTCAGACAAC AATAATCATT TATTATCCAT CATGGTTTTT GTGGGCTAGA 16620

AATTCAGATA GGGTACAGTG GGGAGGGTTC ACCCTGCTCC ACAGCATCTG GAAGATCTGA 16680

- 20 (2) INFORMATION FOR SEQ ID NO: 62:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 50 base pairs
    - (B) TYPE: nucleic acid
- 25 (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA
- 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 62:

ATGCGCGCCG GCAGCCAACA TGCTGCTGCT GCTGCTGGTG CCTCTCTTCC

50

(2) INFORMATION FOR SEQ ID NO: 63:

35

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 25 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
- 0 (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63: GGTCATCCAG CGTTGAGGTG AAGAC 25 (2) INFORMATION FOR SEQ ID NO: 64: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs 10 (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA 15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 64: GAAGGTTGCC AGATTATACA TCCGC 25 20 (2) INFORMATION FOR SEQ ID NO: 65: (i) SEQUENCE CHARACTERISTICS: 25 (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 65: CCACGATGAT TCGAGCATCT TGACG 25 35 (2) INFORMATION FOR SEQ ID NO: 66: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear

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(ii) MOLECU	JLE TYPE: DNA		å		
(xi) SEQUE	NCE DESCRIPTION:	SEQ ID NO:	66:		
CTGGTTCCTC CCA	ATGTG				18
(2) INFORMATION	N FOR SEQ ID NO:	67:			
(i) CEOUEN	IOE CUADACMEDICMI	00.			
	NCE CHARACTERISTI LENGTH: 21 base p				
	MANGIH: 21 base p MYPE: nucleic aci				
	STRANDEDNESS: sin				
	COPOLOGY: linear	gie			
(ii) MOLECU	JLE TYPE: DNA				
(xi) SEQUEN	NCE DESCRIPTION:	SEQ ID NO:	67:		
CCAGTGGACT ATGA	AGATTGA G				21
(2) INFORMATION	FOR SEQ ID NO:	68:			
	ICE CHARACTERISTI				
	ENGTH: 18 base p				
	TYPE: nucleic aci				
	STRANDEDNESS: sin COPOLOGY: linear	gie .			
(ii) MOLECU	JLE TYPE: DNA				•
(xi) SEQUEN	ICE DESCRIPTION:	SEQ ID NO:	68:		
CTGGTTCCTC CCAP	ATGTG				18
(2) INFORMATION	FOR SEQ ID NO:	69:			
(i) SEQUEN	CE CHARACTERISTI	CS:		_	
(A) I	ENGTH: 21 base p	airs		•	
(B) T	TYPE: nucleic aci	đ			
H1865-1 WO SEC	2	121	<del>.</del>		<u> </u>
	-				

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 69:

CCAGTGGACT ATGAGATTGA G

21

10

15

(2) INFORMATION FOR SEQ ID NO: 70:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2700 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA to mRNA

20

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

25 (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

(ix) FEATURE:

(A) NAME/KEY: CDS

30 (B) LOCATION: 1..2700

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 70:

ATG TTG CTG CTG CTA CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC 48

35 Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly

1 5 10 15

GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC 96

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile

40 20 25 30

ATA CAC CCG CCC TGG GAA GGG GGC ATC AGG TAC CGG GGC CTG ACT CGG 144

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg

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GAC CAG GTG AAG GCT ATC AAC TTC CTG CCA GTG GAC TAT GAG ATT GAG Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu TAT GTG TGC CGG GGG GAG CGC GAG GTG GTG GGG CCC AAG GTC CGC AAG Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys TGC CTG GCC AAC GGC TCC TGG ACA GAT ATG GAC ACA CCC AGC CGC TGT Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys GTG AAT CGA ACG CCA CAC TCA GAA CGG CGC GCA GTG TAC ATC GGG GCA Val Asn Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly Ala CTG TTT CCC ATG AGC GGG GGC TGG CCA GGG GGC CAG GCC TGC CAG CCC Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln Pro GCG GTG GAG ATG GCG CTG GAG GAC GTG AAT AGC CGC AGG GAC ATC CTG Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile Leu CCG GAC TAT GAG CTC AAG CTC ATC CAC CAC GAC AGC AAG TGT GAT CCA Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp Pro GGC CAA GCC ACC AAG TAC CTA TAT GAG CTG CTC TAC AAC GAC CCT ATC Gly Gln Ala Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile AAG ATC ATC CTT ATG CCT GGC TGC AGC TCT GTC TCC ACG CTG GTG GCT Lys Ile Ile Leu Met Pro Gly Cys Ser Ser Val Ser Thr Leu Val Ala GAG GCT GCT AGG ATG TGG AAC CTC ATT GTG CTT TCC TAT GGC TCC AGC Glu Ala Ala Arg Met Trp Asn Leu Ile Val Leu Ser Tyr Gly Ser Ser TCA CCA GCC CTG TCA AAC CGG CAG CGT TTC CCC ACT TTC TTC CGA ACG 

	Ser	Pro	Ala	Leu	Ser	Asn	Arg	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	
		210					215					220					
	CAC	CCA	<b>ጥ</b> ር እ	GCC	aca	רייר	CAC	አአሮ	CCT	»cc	ccc	cmc	222	CTTC	ттт		720
5															Phe		720
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	AAG	TGG	GGC	TGG	AAG	AAG	ATT	GCT	ACC	ATC	CAG	CAG	ACC	ACT	GAG	GTC	768
	Lys	Trp	Gly	Trp	Lys	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	Val	
10					245					250					255		
	ሙጥር	ልርጥ	ጥሮር	ልሮጥ	CTG	GAC	GAC	Cutc	CAC	CAA	CCA	C/T/C	አአር	GVG	GCT	CCA	816
															Ala		910
				260		•	•		265		3			270			
15																	
	ATT	GAG	ATT	ACT	TTC	CGC	CAG	AGT	TTC	TTC	TCA	GAT	CCA	GCT	GTG	CCC	864
	Ile	Glu		Thr	Phe	Arg	Gln		Phe	Phe	Ser	Asp		Ala	Val	Pro	
			275					280					285				
20	GTC	AAA	AAC	CTG	AAG	CGC	CAG	GAT	GCC	CGA	ልጥሮ	ልጥሮ	GTG.	GGA	CTT	ጥጥር	912
															Leu		712
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25		Glu	Thr	Glu	Ala		Lys	Val	Phe	Cys		Val	Tyr	Lys	Glu	_	
	305					310					315					320	
	CTC	TTT	GGG	AAG	AAG	TAC	GTC	TGG	TTC	CTC	АТТ	GGG	TGG	TAT	GCT	GAC	1008
															Ala		
30					325					330					335	•	
															GAT		1056
	ASN	Trp	Pne	Lys 340	TIE	ıyr	Asp	Pro	345	TIE	Asn	Cys	Thr	350	Asp	Glu	
35				240					343					330			
•	ATG	ACT	GAG	GCG	GTG	GAG	GGC	CAC	ATC	ACA	ACT	GAG	АТТ	GTC	ATG	CTG	1104
	Met	Thr	Glu	Ala	Val	Glu	Gly	His	Ile	Thr	Thr	Glu	Ile	Val	Met	Leu	
			355					360					365				
40															GAA		1152 <sub>v</sub>
	Asn		Ala	Asn	Tnr	Arg	375	тте	ser	Asn	Met		Ser	GIn	Glu	rne	
		370					3/3					380					

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	GTG	GAG	AAA	СТА	ACC	AAG	CGA	CTG	AAA	AGA	CAC	CCT	GAG	GAĠ	ACA	GGA	1200
	Val	Glu	Lys	Leu	Thr	Lys	Arg	Leu	Lys	Arg	His	Pro	Glu	Glu	Thr	Gly	
	385					390					395					400 .	
5	GGC	TTC	CAG	GAG	GCA	CCG	CTG	GCC	TAT	GAT	GCC	ATC	TGG	GCC	TTG	GCA	1248
	Gly	Phe	Gln	Glu	Ala	Pro	Leu	Ala	Tyr	Asp	Ala	Ile	Trp	Ala	Leu	Ala	
					405					410					415		
											GGC						1296
10	Leu	Ala	Leu	420	ьys	Thr	ser	GIA	425	GIA	Gly	Arg	ser	<del>-</del>	val	Arg	
				420					423					430			
	CTG	GAG	GAC	TTC	AAC	TAC	AAC	AAC	CAG	ACC	АТТ	ACC	GAC	CAA	ATC	TAC	1344
											Ile						
15			435			=		440					445			•	
	CGG	GCA	ATG	AAC	TCT	TCG	TCC	TTT	GAG	GGT	GTC	TCT	GGC	CAT	GTG	GTG	1392
	Arg	Ala	Met	Asn	Ser	Ser	Ser	Phe	Glu	Gly	Val	Ser	Gly	His	Val	Val	
		450					455					460					
20																	
					_						ACG						1440
		Asp	Ala	Ser	СТĀ		Arg	Met	Ala	Trp	Thr	Leu	Ile	Glu	Gln		
	465					470					475					480	
25	CAG	GGT	GGC	AGC	TAC	AAG	AAG	ATT	GGC	TAC	ТАТ	GAC	AGC	ACC	AAG	GAT	1488
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	GAT	CTT	TCC	TGG	TCC	AAA	ACA	GAT	AAA	TGG	АТТ	GGA	GGG	TCC	ccc	CCA	1536
30	Asp	Leu	Ser	Trp	Ser	Lys	Thr	Asp	Lys	Trp	Ile	Gly	Gly	Ser	Pro	Pro .	
				500					505					510		•	
											•						
											CGC						1584
	Ala	Asp		Thr	Leu	Val	Ile	_	Thr	Phe	Arg	Phe		Ser	Gln	Lys	
35			515					520					525				
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	GTT	GTC	TGT	CTG	TCC	TTT	AAC	ATC	TAC	AAC	TCA	CAT	GTC	CGT	TAT	ATC	1680
	Val	Val	Cys	Leu	Ser	Phe	Asn	Ile	Tyr	Asn	Ser	His	Val	Arg	Tyr	Ile	
	545					550					555					560	
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A	sn	Thr	Trp	Leu	Gly	Ile	Phe	Tyr	Gly	Tyr	Lys	Gly	Leu	Leu	Leu	Leu	
A	LAT	ACA	TGG	СТТ	GGC	ATT	TTC	TAT	GGT	TAC	AAG	GGG	CTG	CTG	CTG	CTG	2160
		690					073					700					
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						CCC											2112
			675					680					685				
н	lis	Arg	Thr	Ile	Glu	Thr	Phe	Ala	Lys	Glu	Glu	Pro	Lys	Glu	Asp	Ile	
30 C	AC	CGG	ACC	АТТ	GAG	ACA	TTT	GCC	AAG	GAG	GAA	CCT	AAG	GAA	GAT	ATT	2064
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					645					650					655		
L	ys	Thr	Leu	Glu	Pro	Trp	Lys	Leu	Tyr	Ala	Thr	Val	Gly	Leu	Leu	Val	
A	AG	ACT	CTG	GAA	ccc	TGG	AAG	CTG	TAT	GCC	ACA	GTG	GGC	CTG	CTG	GTG	1968
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						TTC Phe											1920
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15 G	ly	Leu	Gly	Phe	Ser	Leu		Tyr	Gly	Ser	Met	Phe	Thr	Lys	Ile	Trp	
G	GC	CTG	GGC	TTT	AGT	CTG	GGC	TAC	GGT	TCC	ATG	TTC	ACC	AAG	ATT	TGG	1872
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				580					585					590			
L	eu	Ala	Leu		Ala	Val	Phe	Pro		Gly	Leu	Asp	Gly		His	Ile	
C'	TG	GCT	TTA	GCT	GCT	GTC	TTC	CCC	CTG	GGG	CTC	GAT	GGT	TAC	CAC	ATT	1776
5																	
					565					570				-	575		
						Asn											
C	'AG	AAC	тса	CAG	CCC	AAC	CTG	AAC	AAC	СТС	АСТ	GCT	GTG	GGC	ጥርር	тса	1728
C	AG	AAC	TC.	A	A CAG	A CAG CCC	A CAG CCC AAC	A CAG CCC AAC CTG	A CAG CCC AAC CTG AAC	A CAG CCC AAC CTG AAC AAC	A CAG CCC AAC CTG AAC AAC CTG	A CAG CCC AAC CTG AAC AAC CTG ACT	A CAG CCC AAC CTG AAC AAC CTG ACT GCT	A CAG CCC AAC CTG AAC AAC CTG ACT GCT GTG	A CAG CCC AAC CTG AAC AAC CTG ACT GCT GTG GGC	A CAG CCC AAC CTG AAC AAC CTG ACT GCT GTG GGC TGC	A CAG CCC AAC CTG AAC AAC CTG ACT GCT GTG GGC TGC TCA

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					725					730			٠		735	• •		-
	ATC	AAT	GAT	CAC	CGG	GCT	GTG	GGC	ATG	GCT	ATC	TAC	AAT	GTG	GCA	GTC	225	6
												Tyr						
5				740					745			-		750				
	CTG	TGC	CTC	ATC	ACT	GCT	CCT	GTC	ACC	ATG	ATT	CTG	TCC	AGC	CAG	CAG	230	4
	Leu	Cys	Leu	Ile	Thr	Ala	Pro	Val	Thr	Met	Ile	Leu	Ser	Ser	Gln	Gln		
			755					760					765					
10																		
	GAT	GCA	GCC	TTT	GCC	TTT	GCC	TCT	CTT	GCC	ATA	GTT	TTC	TCC	TCC	TAT	235	2
	Asp	Ala	Ala	Phe	Ala	Phe	Ala	Ser	Leu	Ala	Ile	Val	Phe	Ser	Ser	Tyr		
		770					775					780						
15	ATC	ACT	СТТ	GTT	GTG	CTC	ттт	GTG	CCC	AAG	ATG	CGC	AGG	CTG	ATC	ACC	240	)
	Ile	Thr	Leu	Val	Val	Leu	Phe	Val	Pro	Lys	Met	Arg	Arg	Leu	Ile	Thr		
	785					790					795					800		
	CGA	GGG	GAA	TGG	CAG	TCG	GAG	GCG	CAG	GAC	ACC	ATG	AAG	ACA	GGG	TCA	2448	3
20	Arg	Gly	Glu	Trp	Gln	Ser	Glu	Ala	Gln	Asp	Thr	Met	Lys	Thr	Gly	Ser		
					805					810					815			
	TCG	ACC	AAC	AAC	AAC	GAG	GAG	GAG	AAG	TCC	CGG	CTG	TTG	GAG	AAG	GAG	2496	5
	Ser	Thr	Asn	Asn	Asn	Glu	Glu	Glu	Lys	Ser	Arg	Leu	Leu	Glu	Lys	Glu		
25				820					825					830				
	AAC	CGT	GAA	CTG	GAA	AAG	ATC	ATT	GCT	GAG	AAA	GAG	GAG	CGT	GTC	TCT	2544	1
	Asn	Arg	Glu	Leu	Glu	Lys	Ile	Ile	Ala	Glu	Lys	Glu	Glu	Arg	Val	Ser		
			835					840					845					
30																		
			CGC									CTC				CGC	2592	!
	Glu		Arg	His	Gln	Leu		Ser	Arg	Gln	Gln	Leu	Arg	Ser	Arg	Arg		
		850					855					860						
35	CAC	CCA	CCG	ACA	ccc	CCA	GAA	ccc	TCT	GGG	GGC	CTG	ccc	AGG	GGA	ccc	2640	)
	His	Pro	Pro	Thr	Pro	Pro	Glu	Pro	Ser	Gly	Gly	Leu	Pro	Arg	Gly	Pro		
	865					870					875					880		
	CCT	GAG	ccc	ccc	GAC	CGG	CTT	AGC	TGT	GAT	GGG	AGT	CGA	GTG	САТ	TTG	2688	}
40	Pro	Glu	Pro	Pro	Asp	Arg	Leu	Ser	Cys	Asp	Gly	Ser	Arg	Val	His	Leu		ı
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SUBSTITUTE SHEET (RULE 26)

Leu Tyr Lys \*

5 (2) INFORMATION FOR SEQ ID NO: 71:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 899 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 71:

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Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly

1 5 10 15

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile 20 25 30

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg

35 40 45

25 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 55 60

Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys
65 70 75 80

Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys
85 90 95

Val Asn Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly Ala 35 100 105 110

Leu Phe Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln Pro 115 120 125

40 Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile Leu 130 135 140

Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp Pro

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Gly Gln Ala Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile Leu Met Pro Gly Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr 215 · 15 His Pro Ser Ala Thr Leu His Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly Trp Lys Lys Ile Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr Ser Thr Leu Asp Asp Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile Thr Phe Arg Gln Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Ile Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe 

Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile 35 Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Asn Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu 

Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Arg Lys Met Asn Thr Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Leu Gly Ile Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu Val Val Leu Phe Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu Ala Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile Ile Ala Glu Lys Glu Glu Arg Val Ser H1865-1 WO SEQ 

835

840

845

Glu Leu Arg His Gln Leu Gln Ser Arg Gln Gln Leu Arg Ser Arg Arg 850 855 860

5

His Pro Pro Thr Pro Pro Glu Pro Ser Gly Gly Leu Pro Arg Gly Pro 865 870 875 880

Pro Glu Pro Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His Leu 10 885 890 895

Leu Tyr Lys

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- (2) INFORMATION FOR SEQ ID NO: 72:
  - (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2518 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA to mRNA

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- (iii) HYPOTHETICAL: NO
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo sapiens

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- (ix) FEATURE:
  - (A) NAME/KEY: CDS
  - (B) LOCATION:1..294

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 72:

ATG TTG CTG CTG CTA CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly

40 1 5 10 15

GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC
Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile

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				TGG												144
Ile	His	Pro 3!		Trp	Glu	Gly	Gly 40		Arg	Tyr	Arg	Gly 45	Leu	Thr	Arg	
GAC	CAG	GTG	AAG	GCT	ATC	AAC	TTC	CTG	CCA	GTG	GAC	TAT	GAG	ATT	GAG	192
Asp		Val	Lys	Ala	Ile		Phe	Leu	Pro	Val		Tyr	Glu	Ile	Glu	
	50					55					60					
				GGG												240
Tyr 65	Val	Cys	Arg	Gly	Glu 70	Arg	Glu	Val	Val	Gly 75	Pro	Lys	Val	Arg	80	
				GGC												288
Cys	Leu	Ala	Asn	Gly 85	Ser	Trp	Thr	Asp	Met 90	Asp	Thr	Pro	Ser	Arg 95	Cys	·
GTG	TGA	TCC	AGGC	CAA C	CCA	CCAAC	GT AC	CTAT	ratg <i>i</i>	A GCT	rgcto	CTAC	AAC	GACC	CTA	344
Val	*															
TCA	AGATO	CAT (	CCTT	ATGCC	CT GO	CTG	CAGCT	CTO	STCTO	CAC	GCTC	GTGC	CT (	GAGG	CTGCTA	. 404
GGA'	rgtgo	GAA (	CCTC	ATTG1	rg ci	TTCC	CTATO	GC1	rccac	CTC	ACCA	AGCCC	CTG :	rcaa)	ACCGGC	464
AGC	FTTTC	cc (	CACT	rtcti	rc co	BAACC	GCACC	CAT	CAGO	CAC	ACTO	CAC	AAC (	CCTAC	CCGCG	524
TGA	AACTO	CTT 1	rgaaj	AAGTO	G GC	CTGC	SAAGA	AG?	ATTGC	TAC	CATO	CAGO	CAG A	ACCAC	TGAGG	. 584
TCT	rcaci	TTC (	GACTO	CTGG#	AC GA	ACCTO	GAGG	AAC	GAGT	GAA	GGAG	GCTG	GA 1	ATTG <i>I</i>	GATTA	644
CTT'	rccgo	CCA (	GAGTT	rtcti	C TO	CAGAT	CCAG	CTC	STGCC	CGT	CAAA	AACC	TG A	AAGCO	CCAGG	704
ATG	CCG/	AAT (	CATCO	STGGG	SA CI	TTTT	TATG	AG/	\CTG#	AGC	CCGG	SAAAG	TT ?	rttt	STGAGG	764
TGT	ACAA	GGA (	GCGT	CTCTI	T GO	GAAG	SAAGT	ACC	STCTO	GTT	CCTC	ATTO	GG 7	rggt <i>i</i>	TGCTG	824
ACA	ATTGO	STT (	CAAGA	ATCTA	AC GA	CCCI	ГТСТА	TC	ACTO	CAC	AGTO	GATO	SAG A	ATGAC	TGAGG	884
CGG'	rggao	GG (	CCAC	ATCAC	CA AC	CTGAC	SATTG	TCA	TGCI	GAA	TCCI	GCCA	LAT A	ACCC(	CAGCA	944
TTT	CCAA	CAT (	GACA	rccc <i>i</i>	AG GA	\ATT1	rgtgg	ag/	AACI	CAAC	CAAG	CGAC	TG A	AAAA	BACACC	. 1004
H18	TTCCAACAT GACATCCC					··· <u>-</u>	1	33								

	CTGAGGAGAC	AGGAGGCTTC	CAGGAGGCAC	CGCTGGCCTA	TGATGCCATC	TGGGCCTTGG	1064
5	CACTGGCCCT	GAACAAGACA	TCTGGAGGAG	GCGGCCGTTC	TGGTGTGCGC	CTGGAGGACT	1124
3	TCAACTACAA	CAACCAGACC	ATTACCGACC	AAATCTACCG	GGCAATGAAC	TCTTCGTCCT	1184
	TTGAGGGTGT	CTCTGGCCAT	GTGGTGTTTG	ATGCCAGCGG	CTCTCGGATG	GCATGGACGC	1244
10	TTATCGAGCA	GCTTCAGGGT	GGCAGCTACA	AGAAGATTGG	CTACTATGAC	AGCACCAAGG	1304
	ATGATCTTTC	CTGGTCCAAA	ACAGATAAAT	GGATTGGAGG	GTCCCCCCA	GCTGACCAGA	1364
15	CCCTGGTCAT	CAAGACATTC	CGCTTCCTGT	CACAGAAACT	CTTTATCTCC	GTCTCAGTTC	1424
נו	TCTCCAGCCT	GGGCATTGTC	CTAGCTGTTG	TCTGTCTGTC	CTTTAACATC	TACAACTCAC	1484
	ATGTCCGTTA	TATCCAGAAC	TCACAGCCCA	ACCTGAACAA	CCTGACTGCT	GTGGGCTGCT	1544
20	CACTGGCTTT	AGCTGCTGTC	TTCCCCCTGG	GGCTCGATGG	TTACCACATT	GGGAGGAACC	1604
	AGTTTCCTTT	CGTCTGCCAG	GCCCGCCTCT	GGCTCCTGGG	CCTGGGCTTT	AGTCTGGGCT	1664
25	ACGGTTCCAT	GTTCACCAAG	ATTTGGTGGG	TCCACACGGT	CTTCACAAAG	AAGGAAGAAA	1724
-5	AGAAGGAGTG	GAGGAAGACT	CTGGAACCCT	GGAAGCTGTA	TGCCACAGTG	GGCCTGCTGG	1784
	TGGGCATGGA	TGTCCTCACT	CTCGCCATCT	GGCAGATCGT	GGACCCTCTG	CACCGGACCA	1844
30	TTGAGACATT	TGCCAAGGAG	GAACCTAAGG	AAGATATTGA	CGTCTCTATT	CTGCCCCAGC .	1904
	TGGAGCATTG	CAGCTCCAGG	AAGATGAATA	CATGGCTTGG	CATTTTCTAT	GGTTACAAGG	1964
35	GGCTGCTGCT	GCTGCTGGGA	ATCTTCCTTG	CTTATGAGAC	CAAGAGTGTG	TCCACTGAGA	2024
33	AGATCAATGA	TCACCGGGCT	GTGGGCATGG	СТАТСТАСАА	TGTGGCAGTC	CTGTGCCTCA	2084
	TCACTGCTCC	TGTCACCATG	ATTCTGTCCA	GCCAGCAGGA	TGCAGCCTTT	GCCTTTGCCT	2144
40	CTCTTGCCAT	AGTTTTCTCC	тсстататса	CTCTTGTTGT	GCTCTTTGTG	CCCAAGATGC	2204
	GCAGGCTGAT	CACCCGAGGG	GAATGGCAGT	CGGAGGCGCA	GGACACCATG	ÂAGACAGGGT	2264

CATCGACCAA CAACAACGAG GAGGAGAAGT CCCGGCTGTT GGAGAAGGAG AACCGTGAAC 2324 TGGAAAAGAT CATTGCTGAG AAAGAGGAGC GTGTCTCTGA ACTGCGCCAT CAACTCCAGT 5 CTCGGCAGCA GCTCCGCTCC CGGCGCCACC CACCGACACC CCCAGAACCC TCTGGGGGCC 2444 TGCCCAGGGG ACCCCCTGAG CCCCCCGACC GGCTTAGCTG TGATGGGAGT CGAGTGCATT 2504 TGCTTTATAA GTGA 2518

10

15

## (2) INFORMATION FOR SEQ ID NO: 73:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 97 amino acids
  - (B) TYPE: amino acid
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 73:

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly 10

25

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile 20 25

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg 40

Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 55

Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys 65 70 75

Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys 85 90

40

Val

(2) INFORMATION FOR SEQ ID NO: 74: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2679 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 10 (ii) MOLECULE TYPE: cDNA to mRNA (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO 15 (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1.. 2679 (xi) SEQUENCE DESCRIPTION: SEO ID NO: 74: ATG TTG CTG CTG CTG CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC 48 Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly 10 GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC 96 Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile 20 ATA CAC CCG CCC TGG GAA GGG GGC ATC AGG TAC CGG GGC CTG ACT CGG 144 Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg 35 35 40 45 GAC CAG GTG AAG GCT ATC AAC TTC CTG CCA GTG GAC TAT GAG ATT GAG 192 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 55 60 40 TAT GTG TGC CGG GGG GAG CGC GAG GTG GTG GGG CCC AAG GTC CGC AAG 240 Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys 70

136

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											<b></b>				<sub>P</sub>		٠	
						Thr												, 50
	CCm	ACC	ልሞር	ርልር	CyG	ACC	ልሮጥ	G7G	ርጥር	ጥጥር	ልሮጥ	ጥርር	ልሮሞ	رس <i>ت</i> ,	GAC	GAC		768
40	225					230					235					240		υ
		Pro	Thr	Arg	Val	Lys	Leu	Phe	Glu	Lys	-	Gly	Trp	Lys	Lys			
						AAA												720
		210					215					220						
35	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	His	Pro	Ser	Ala	Thr	Leu	His		
	CAG	CGT	TTC	CCC	ACT	TTC	TTC	CGA	ACG	CAC	CCA	TCA	GCC	ACA	CTC	CAC		672
			195				=	200					205			-		
=						Tyr												<b>-</b>
30	CTC	АТТ	GTG	CTT	TCC	TAT	GGC	TCC	AGC	TCA	CCA	GCC	CTG	TCA	AAC	CGG		624
				190					185					190				
	Cys	Ser	Ser	Val 180	Ser	Thr	Leu	val		GIu	Ala	Ala	Arg		Trp	Asn		
						ACG												576
25																		
					165					170					175			
	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro	Ile	Lys	Ile	Ile	Leu	Met	Pro	Gly		
	TAT	GAG	CTG	CTC	TAC	AAC	GAC	CCT	ATC	AAG	ATC	ATÇ	СТТ	ATG	ССТ	GGC		528
																•		
20	145			<i>ت</i> است.		150	-,,0			~~,	155	*********	****	Lys	-1-	160		
						Lys												400
	Δηγ	ראַר	CAC	GAC	ልሮሮ	AAG	ጥርጥ	ርልጥ	ርር እ	GGC	ר א מ	GCC	እሮር	מממ	መአር	СШУ		480
		130					135					140						
15	Asp		Asn	Ser	Arg	Arg		Ile	Leu	Pro	Asp		Glu	Leu	Lys	Leu		
						AGG												432
			115					120					125					
						Ala												
10	TGG	CCA	GGG	GGC	CAG	GCC	TGC	CAG	ccc	GCG	GTG	GAG	ATG	GCG	CTG	GAG		384
				100					103					110				
	GIU	nr 9	nt 9	100	vaı	TYL	116	ату	105	neu	rne	PIO	met	110	GIĀ	GTÅ		
						TAC Tyr												336
5	CAA	ccc	ccc	CCA	CMC	መልር	איייט	ccc	CCX	CITIC	Man.	000	a mc	3.00		666		226
_					85					90					95			
	Cys	Leu	Ala	Asn		Ser	Trp	Thr	Asp		Asp	Thr	Pro	Ser	Arg	Суѕ	•	
						TCC												288
	•											•	•	•	•	• • •		-

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					245					250		٠	•	••	255	• • •	· <del>-</del>
	omo.	03.0	C2.2	CC.	CEC	220	CAC	CCM	CCA	» mm	CAC	3.00	3.00	mmo	000	<b>616</b>	016
															CGC		816
5	Deu	Giu	GIU	260	Vai	Lys	GIU	A1G	265	116	Giu	110	1111	270	ALG	GIII	
•				200					203								
	AGT	TTC	TTC	TCA	GAT	CCA	GCT	GTG	ccc	GTC	AAA	AAC	CTG	AAG	CGC	CAG	864
	Ser	Phe	Phe	Ser	Asp	Pro	Ala	Val	Pro	Val	Lys	Asn	Leu	Lys	Arg	Gln	
			275					280					285				
10																	
	GAT	GCC	CGA	ATC	ATC	GTG	GGA	CTT	TTC	ТАТ	GAG	ACT	GAA	GCC	CGG	AAA	912
	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	Phe	Tyr	Glu	Thr	Glu	Ala	Arg	Lys	
		290					295					300					
											,						
15															TAC		960
		Phe	Cys	Glu	Val		Lys	Glu	Arg	Leu		Gly	Lys	Lys	Tyr		
	305					310					315					320	
	тсс	መጥር	ርሞር	ል ጥጥ	ccc	тас	ጥልጥ	CCT	GAC	አልጥ	таа	ሙሙር	ልልር	ልጥሮ	TAC	GAC	1008
20															Tyr		1008
20					325	1-6	- 1 -			330			-,,		335		
										,							
	ССТ	TCT	ATC	AAC	TGC	ACA	GTG	GAT	GAG	ATG	ACT	GAG	GCG	GTG	GAG	GGC	1056
	Pro	Ser	Ile	Asn	Cys	Thr	Val	Asp	Glu	Met	Thr	Glu	Ala	Val	Glu	Gly	
25				340					345					350			
	CAC	ATC	ACA	ACT	GAG	ATT	GTC	ATG	CTG	AAT	CCT	GCC	AAT	ACC	CGC	AGC	1104
	His	Ile		Thr	Glu	Ile	Val		Leu	Asn	Pro	Ala	Asn	Thr	Arg	Ser	
			355					360					365				
30						maa	~~~	~~~	mmm	omo	~~~		am.			~~.	1150
	ATT	TCC		ATG		TCC	CAG		TTT		GAG		CTA	ACC	AAG	CGA	1152
	TTE	370	ASII	met	THE	ser	375	GIU	Pne	vai	GIU	380	rea	THE	Lys	Arg	
		570					373					300					
35	CTG	AAA	AGA	CAC	ССТ	GAG	GAG	ACA	GGA	GGC	TTC	CAG	GAG	GCA	CCG	CTG	1200
55															Pro		
	385	=				390			_	-	395					400	
	GCC	TAT	GAT	GCC	ATC	TGG	GCC	TTG	GCA	CTG	GCC	CTG	AAC	AAG	ACA	TCT	1248
40	Ala	Tyr	Asp	Ala	Ile	Trp	Ala	Leu	Ala	Leu	Ala	Leu	Asn	Lys	Thr	Ser	Ų
					405					410					415		
														•			
	GGA	GGA	GGC	GGC	CGT	TCT	GGT	GTG	CGC	CTG	GAG	GAC	TTC	AAC	TAC	AAC	1296
																· · · · · · · · · · · · · · · · · · ·	
	H18	65-1	wo	SEQ				1	138								

	Gly	Gly	Gly	Gly 420	Arg	Ser	Gly	Val	Arg 425	Leu	Glu	Asp	Phe	Asn 430	Тут	Aśn	
	AAC	CAG	ACC	ATT	ACC	GAC	CAA	ATC	TAC	CGG	GCA	ATG	AAC	TCT	TCG	TCC	1344
5	Asn	Gln		Ile	Thr	Asp	Gln		Tyr	Arg	Ala	Met		Ser	Ser	Ser	
			435					440					445				
	TTT	GAG	GGT	GTC	TCT	GGC	CAT	GTG	GTG	ттт	GAT	GCC	AGC	GGC	тст	CGG	1392
	Phe	Glu	Gly	Val	Ser	Gly	His	Val	Val	Phe	Asp	Ala	Ser	Gly	Ser	Arg	
10		450					455					460					
															AAG		1440
	465	Ala	тр	THE	rea	470	GIU	GIN	rea	GIN	475	GTĀ	Ser	туг	Lys	Lys 480	
15	402					470					4,5					400	
	ATT	GGC	TAC	тат	GAC	AGC	ACC	AAG	GAT	GAT	CTT	TCC	TGG	TCC	AAA	ACA	1488
	Ile	Gly	Tyr	Tyr	Asp	Ser	Thr	Lys	Asp	Asp	Leu	Ser	Trp	Ser	Lys	Thr	
					485					490					495		
20															GTC		1536
	Asp	гÀг	Trp	500	GIÀ	GTA	ser	Pro	505	Ala	Asp	GIN	Thr	510	Val	iie	
				300										310			
	AAG	ACA	TTC	CGC	TTC	CTG	TCA	CAG	AAA	CTC	TTT	ATC	TCC	GTC	TCA	GTT	1584
25	Lys	Thr	Phe	Arg	Phe	Leu	Ser	Gln	Lys	Leu	Phe	Ile	Ser	Val	Ser	Val	
			515					520					525				
															TTT		1632
30	Leu	530	ser	reu	GIY	116	535	neu	Ala	vai	vai	540	rea	ser	Phe	ASII	
30		550					555					240					
	ATC	TAC	AAC	TCA	САТ	GTC	CGT	тат	ATC	CAG	AAC	TCA	CAG	ccc	AAC	CTG	1680
	Ile	Tyr	Asn	Ser	His	Val	Arg	Tyr	Ile	Gln	Asn	Ser	Gln	Pro	Asn	Leu	
	545					550					555					560	
35																	1700
															GTC		1728
	ASN	Asn	Leu	Thr	565	AST	GIA	cys	ser	570	Ala	Leu	Ala	Ald	Val 575	File	
					J. J. J					2,0					5,5		
40	CCC	CTG	GGG	CTC	GAT	GGT	TAC	CAC	ATT	GGG	AGG	AAC	CAG	TTT	CCT	TTC	1776 <sub>0</sub>
	Pro	Leu	Gly	Leu	Asp	Gly	Tyr	His	Ile	Gly	Arg	Asn	Gln	Phe	Pro	Phe	
				580					585					590	•		

								•	•				•				
	GTC	TGC	CAG	GCC	CGC	CTC	TGG	CTC	CTG	GGC	CTG	GGC	ТŤТ	AGT	CTG	GGC	1824
	Val	Cys	Gln	Ala	Arg	Leu	Trp	Leu	Leu	Gly	Leu	Gly	Phe	Ser	Leu	Gly	
			595					600					605				
5	TAC	GGT	TCC	ATG	TTC	ACC	AAG	ATT	TGG	TGG	GTC	CAC	ACG	GTC	TTC	ACA	1872
	Tyr	Gly	Ser	Met	Phe	Thr	Lys	Ile	Trp	Trp	Val	His	Thr	Val	Phe	Thr	
		610					615					620					
		•															
	AAG	AAG	GAA	GAA	AAG	AAG	GAG	TGG	AGG	AAG	ACT	CTG	GAA	ccc	TGG	AAG	1920
10	Lys	Lys	Glu	Glu	Lys	Lys	Glu	Trp	Arg	Lys	Thr	Leu	Glu	Pro	Trp	Lys	
	625					630					635					640	
	CTG	TAT	GCC	ACA	GTG	GGC	CTG	CTG	GTG	GGC	ATG	GAT	GTC	CTC	ACT	CTC	1968
	Leu	Tyr	Ala	Thr	Val	Gly	Leu	Leu	Val	Gly	Met	Asp	Val	Leu	Thr	Leu	
15					645					650					655		
	GCC	ATC	TGG	CAG	ATC	GTG	GAC	CCT	CTG	CAC	CGG	ACC	ATT	GAG	ACA	TTT	2016
	Ala	Ile	$\mathtt{Trp}$	Gln	Ile	Val	Asp	Pro	Leu	His	Arg	Thr	Ile	Glu	Thr	Phe	
				660					665					670			
20																	
	GCC	AAG	GAG	GAA	CCT	AAG	GAA	GAT	ATT	GAC	GTC	TCT	ATT	CTG	CCC	CAG	2064
	Ala	Lys	Glu	Glu	Pro	Гуs	Glu	Asp	Ile	Asp	Val	Ser	Ile	Leu	Pro	Gln	
			675					680					685				
							•										
25								AAG									2112
	Leu		His	Суѕ	Ser	Ser		Lys	Met	Asn	Thr	Trp	Leu	Gly	Ile	Phe	
		690					695					700					
		_			_			CTG									2160
30		GIA	Tyr	rys	GIA		Leu	Leu	Leu	Leu		Ile	Phe	Leu	Ala		
	705					710					715					720	
		300		3 C M	ama	maa	N C/M	a.c		3.00		<b>63.</b> m	~ ~				2222
								GAG									2208
	GIU	THE	гуз	ser	•	ser	THE	Glu	гуз	730	Asn	Asp	HIS	Arg		vai	
35					725					/30					735		
	ccc	λTC	CCT	איייר	TAC.	יתממ	CTC	GCA	CTC	СТС	TCC	CTC	አመሮ	እርጥ	CCM	CCT	2256
																	2230
	GIA	rie C	uta	740	TAT	USII	AGT	Ala	745	±-cu	-ys	₽eu	TIE	750	via	FIU	
40				, · <u>14</u> U					/923					, 50			
40	CTC	»CC	ልጥር	<u>ን</u> ጥጥ	CTG	ጥርር	AGC	CAG	CAG	ጥፋጋ	GC A	GCC	ተመተ	GCC	יטואלי	GCC	2304
								Gln									2304
	vaı	TIIT	755	T16	₽€U	Jer	JEL	760	GAII	wah	770	vra	765	VIG	FIIE	210	
			133					, 00					100				

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												Leu					2332
	<b>J</b> C1	770			*		775	501	-3-		****	780	Val	Val	neu	rne	
5		,,,					.,,					700					
	GTG	ccc	AAG	ATG	CGC	AGG	CTG	ATC	ACC	CGA	GGG	GAA	TGG	CAG	TCG	GAG	2400
	Val	Pro	Lys	Met	Arg	Arg	Leu	Ile	Thr	Arg	Gly	Glu	Trp	Gln	Ser	Glu	
	785					790					795					800	
10	GCG	CAG	GAC	ACC	ATG	AAG	ACA	GGG	TCA	TCG	ACC	AAC	AAC	AAC	GAG	GAG	2448
	Ala	Gln	Asp	Thr	Met	Lys	Thr	Gly	Ser	Ser	Thr	Asn	Asn	Asn	Glu	Glu	
					805					810					815		
	GAG	AAG	TCC	CGG	CTG	TTG	GAG	AAG	GAG	AAC	CGT	GAA	CTG	GAA	AAG	ATC	2496
15	Glu	Lys	Ser	Arg	Leu	Leu	Glu	Lys	Glu	Asn	Arg	Glu	Leu	Glu	Lys	Ile	
				820					825					830			
	ATT	GCT	GAG	AAA	GAG	GAG	CGT	GTC	TCT	GAA	CTG	CGC	CAT	CAA	CTC	CAG	2544
	Ile	Ala	Glu	Lys	Glu	Glu	Arg	Val	Ser	Glu	Leu	Arg	His	Gln	Leu	Gln	
20			835					840					845				
	TCT	CGG	CAG	CAG	CTC	CGC	TCC	CGG	CGC	CAC	CCA	CCG	ACA	CCC	CCA	GAA	2592
	Ser	Arg	Gln	Gln	Leu	Arg	Ser	Arg	Arg	His	Pro	Pro	Thr	Pro	Pro	Glu	
		850					855					860					
25																	
	CCC	TCT	GGG	GGC	CTG	CCC	AGG	GGA	CCC	CCT	GAG	CCC	CCC	GAC	CGG	CTT	2640
	Pro	Ser	Gly	Gly	Leu	Pro	Arg	Gly	Pro	Pro	Glu	Pro	Pro	Asp	Arg	Leu	
	865					870					875					880	
30	AGC	TGT	GAT	GGG	AGT	CGA	GTG	CAT	TTG	CTT	TAT	AAG	TGA				2679
	Ser	Cys	Asp	Gly	Ser	Arg	Val	His	Leu	Leu	Tyr	Lys	*				
					885					890							

- (2) INFORMATION FOR SEQ ID NO: 75:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 892 amino acids
    - (B) TYPE: amino acid
- 40 (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 75:

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly

1 5 10 15

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile
20 25 30

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg
35 40 45

Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 55 60

Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys
65 70 75 80

Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys
85 90 95

Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe Pro Met Ser Gly Gly
100 105 110

Trp Pro Gly Gly Gln Ala Cys Gln Pro Ala Val Glu Met Ala Leu Glu 25 115 120 125

Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu 130 135 140

30 Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala Thr Lys Tyr Leu 145 150 155 160

Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile Leu Met Pro Gly 165 170 175

Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn 180 185 190

Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala Leu Ser Asn Arg

195 200 205

Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser Ala Thr Leu His 210 215 220

35

	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe	Glu	Lys	Trp	Gly	Trp	Lys	Lys	Ile
	225					230					235					240
5	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	Val	Phe	Thr	Ser	Thr	Leu	Asp	Asp
					245					250					255	-
	Lou	C1	C1	λ ~~	17-1	Tura	C1	212	C111	T10	C1	T10	mb ~	Dho	λ <b></b>	C1-
	reu	GIU	GIU		Val	гуз	GIU	MIA		TTE	GIU	TIE	1111		Arg	GIN
				260					265					270		
10																
	Ser	Phe	Phe	Ser	Asp	Pro	Ala	Val	Pro	Val	Lys	Asn	Leu	Lys	Arg	Gln
			275					280					285			
	Asp	Ala	Arg	Ile	Ile	Val	Gly	Leu	Phe	Tyr	Glu	Thr	Glu	Ala	Arg	Lys
15		290					295					300				
	Val	Phe	Cys	Glu	Val	Tyr	Lys	Glu	Arg	Leu	Phe	Gly	Lys	Lys	Tyr	Val
	305		-			310	-		_		315	-	-	-	-	320
20	Trn.	Dhe	Len	Tle	Gly	mrn.	ጥረድ	Δ1 =	Δen	λen	ሞፖካ	Dho	Lare	T10	The same	Aen
20	IID	rne	nea	116	_	пр	ıyı	nia	rsp			rne	כעם	116		vəħ
					325					330					335	
	_	_		_	_			_			_,					
	Pro	Ser	He		Cys	Thr	Val	Asp		Met	Thr	GIU	Ala		Glu	GIĀ
				340					345					350		
25																
	His	Ile	Thr	Thr	Glu	Ile	Val	Met	Leu	Asn	Pro	Ala	Asn	Thr	Arg	Ser
			355					360					365			
	Ile	Ser	Asn	Met	Thr	Ser	Gln	Glu	Phe	Val	Glu	Lys	Leu	Thr	Lys	Arg
30		370					375					380				
	Leu	Lvs	Ara	His	Pro	Glu	Glu	Thr	Glv	Glv	Phe	Gln	Glu	Ala	Pro	Leu
	385	_	<b>3</b>								395					400
	505					330					333					400
	.1.	<b>(</b>	<b>.</b>	31.	<b>71</b> -	M		r			<b>71</b> -	T	<b>&gt;</b>	*	m\	C
35	Ala	Tyr	Asp	Ala	Ile	Trp	Ala	Leu	Ala		AIA	Leu	ASI	ьys		Ser
					405					410					415	
	Gly	Gly	Gly	Gly	Arg	Ser	Gly	Val	Arg	Leu	Glu	Asp	Phe	Asn	Tyr	Asn
				420					425				•	430		
40																
	Asn	Gln	Thr	Ile	Thr	Asp	Gln	Ile	Tyr	Arg	Ala	Met	Asn	Ser	Ser	Ser
			435					440					445	•	•	

Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Asn Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln H1865-1 WO SEQ 

675 680 685

Leu Glu His Cys Ser Ser Arg Lys Met Asn Thr Trp Leu Gly Ile Phe 690 695 700

Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Gly Ile Phe Leu Ala Tyr
705 710 715 720

Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp His Arg Ala Val
725 730 735

Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu Ile Thr Ala Pro 740 745 750

Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala Phe Ala Phe Ala 755 760 765

Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu Val Val Leu Phe 770 775 780

Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu
785 790 795 800

Ala Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu
25 805 810 815

Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile 820 825 830

30 Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu Arg His Gln Leu Gln 835 840 845

Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro Pro Thr Pro Pro Glu 850 855 860

Pro Ser Gly Gly Leu Pro Arg Gly Pro Pro Glu Pro Pro Asp Arg Leu 865 870 875 880

Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr Lys 885 890

(2) INFORMATION FOR SEQ ID NO: 76:

5

35

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5		(i	() ()	A) L: B) T: C) S:	engti YPE :   Trani	HARAGH: 2: nuc: DEDN: OGY:	661   leic ESS:	acie sin	pai: d	rs								
		(ii	) MO	LECU	LE T	YPE:	CDN	A to	mRN	A								
10		(iii)	) <b>HY</b>	POTH	ETIC	AL: 1	NO											
		(iv	) AN	ri-si	ense	: NO												
		(vi	OR:	IGIN	AL S	OURC	Ε:											
15			(2	A) 01	RGAN:	ISM:	Homo	sa <sub>l</sub>	piens	S								
		(ix	) FE															
			-			KEY: ION::		561										
20												_						
		(X1,	) SE(	QUEN(	JE DI	ESCR:	IPTI	ON:	SEQ .	TD NO	): /(	o:						
						CTA										-		48
25	met 1	ren	Leu	ьeu	Leu 5	Leu	Leu	Ala	Pro	10	Pne	rea	Arg	Pro	15	GIĀ		
	ccc	ccc	ccc	ccc	CAC	ACC	ccc	220	ccc	» CC	<b>ጥ</b> ር እ	CAA	CCT	mc c	CAC	አመሮ		96
						Thr												90
20				20					25					30				
30	ATA	CAC	CCG	ccc	TGG	GAA	GGG	GGC	ATC	AGG	TAC	CGG	GGC	CTG	ACT	CGG		144
	Ile	His		Pro	Trp	Glu	Gly	_	Ile	Arg	Tyr	Arg	_	Leu	Thr	Arg		
			35					40					45					
35	GAC	CAG	GTG	AAG	GCT	ATC	AAC	TTC	CTG	CCA	GTG	GAC	TAT	GAG	ATT	GAG		192
	Asp	Gln 50	Val	Lys	Ala	Ile	Asn 55	Phe	Leu	Pro	Val	Asp	Tyr	Glu	Ile	Glu		
		50					33					00						
						GAG												240
40	_	Val	Cys	Arg	Gly	Glu 70	Arg	Glu	Val	Val		Pro	Lys	Val	Arg	Lys 80		ı
	65					, 0					75				•	30		
	TGC	CTG	GCC	AAC	GGC	TCC	TGG	ACA	GAT	ATG	GAC	ACA	CCC	AGC	CGC	TGT	•	288
	H18	65-1	wo	SEQ					146							<u>.</u>		

	Cys	Leu	Ala	Asn	Gly 85	Ser	Trp	Thr	qaA	Met 90	Asp	Thr	Pro	Ser	Arg 95	Cys	
	GTG	ААТ	CGA	ACG	CCA	CAC	TCA	GAA	CGG	CGC	GCA	GTG	TAC	ATC	GGG	GCA	336
5	Val	Asn	Arg	Thr	Pro	His	Ser	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	Ala	
				100					105					110			
	CTG	TTT	ccc	GCG	GTG	GAG	ATG	GCG	CTG	GAG	GAC	GTG	AAT	AGC	CGC	AGG	384
	Leu	Phe	Pro	Ala	Val	Glu	Met	Ala	Leu	Glu	Asp	Val	Asn	Ser	Arg	Arg	
10			115					120					125				
	GAC	ATC	CTG	CCG	GAC	TAT	GAG	CTC	AAG	CTC	ATC	CAC	CAC	GAC	AGC	AAG	432
	Asp	Ile	Leu	Pro	Asp	Tyr	Glu	Leu	Lys	Leu	Ile	His	His	Asp	Ser	Lys	
		130					135					140					
15																	
						GCC											480
	_	Asp	Pro	Gly	Gln	Ala	Thr	Lys	Tyr	Leu		Glu	Leu	Leu	Tyr		
	145					150					155					160	
											<b></b>						500
20						ATC				_				_			528
	Asp	Pro	IIe	гÀг	11e	Ile	Leu	Met	Pro	170	Суѕ	ser	ser	vaı	5er 175	Tnr	
					100					170					1/5		
	СПС	GTG	GCT	GAG	GCT	GCT	AGG	ATG	TGG	AAC	СТС	АТТ	GTG	СТТ	TCC	тат	576
25						Ala						_					
				180			5		185					190			
	GGC	TCC	AGC	TCA	CCA	GCC	CTG	TCA	AAC	CGG	CAG	CGT	TTC	ccc	ACT	TTC	624
	Gly	Ser	Ser	Ser	Pro	Ala	Leu	Ser	Asn	Arg	Gln	Arg	Phe	Pro	Thr	Phe	
30			195					200					205				
	TTC	CGA	ACG	CAC	CCA	TCA	GCC	ACA	CTC	CAC	AAC	CCT	ACC	CGC	GTG	AAA	672
	Phe	Arg	Thr	His	Pro	Ser	Ala	Thr	Leu	His	Asn	Pro	Thr	Arg	Val	Lys	
		210					215					220					
35																	
	CTC	TTT	GAA	AAG	TGG	GGC	TGG	AAG	AAG	ATT	GCT	ACC	ATC	CAG	CAG	ACC	720
	Leu	Phe	Glu	Lys	Trp	Gly	Trp	Lys	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr	
	225					230					235					240	
																	<b></b>
40						TCG											768 <sub>v</sub>
	Thr	Glu	Val	Phe		Ser	Thr	Leu	Asp		Leu	Glu	Glu	Arg		Ĺys	
					245					250					255		

				<u> </u>					1.40	_								—
				420					425					430				
	Gly	Val	Arg		Glu	Asp	Phe	Asn		Asn	Asn	Gln	Thr		Thr	Asp		
						GAC											. 12	96
40																		ι
					405					410					415			
	Ala	Leu	Ala	Leu	Ala	Leu	Asn	Lys	Thr	Ser	Gly	Gly	Gly	Gly	Arg	Ser		
	GCC	TTG	GCA	CTG	GCC	CTG	AAC	AAG	ACA	TCT	GGA	GGA	GGC	GGC	CGT	TCT	12	48
35	385		~ <b>.</b> y	OLY		390	u				395	-1-	بر د			400		
						Gln											14	55
	CAC	» C	CCA	ccc	mma	CAG	CAC	CCZ	ccc	CTVC	CCC	መጸመ	CVD	ccc	አመር	TVCC:	10	00
		370					375					380						
30	Gln		Phe	Val	Glu	Lys	Leu	Thr	Lys	Arg	Leu		Arg	His	Pro	Glu		
	CAG	GAA	TTT	GTG	GAG	AAA	CTA	ACC	AAG	CGA	CTG	AAA	AGA	CAC	CCT	GAG	11	.52
			355					360					365					
						Ala		_			_				_		- <b>-</b>	-
25	GTC	ATG	CTG	AAT	CCT	GCC	ААТ	ACC	CGC	AGC	ATT	TCC	AAC	ATG	ACA	TCC	11	.04
				340					345					350				
	Val	Asp	Glu		Thr	Glu	Ala	Val		Gly	His	Ile	Thr		Glu	Ile		
						GAG											10	56
20																		
					325					330					335			
	Tyr	Ala	Asp	Asn	Trp	Phe	Lys	Ile	Tyr	Asp	Pro	Ser	Ile	Asn	Cys	Thr		
	ТАТ	GCT	GAC	AAT	TGG	TTC	AAG	ATC	TAC	GAC	CCT	TCT	ATC	AAC	TGC	ACA	10	08
13	203					310					J 4. J					520		
15	ьуs 305	GIU	wrg	ьeu	rne	Gly 310	гуѕ	гÃ2	ıyı	val	315	rne	neu	116	стЛ	320		
						GGG				_		_		_	_		9	60
						000			mr -	a==	mee	m=~				<b>ma</b> -	_	
		290					295					300						
10	Gly	Leu	Phe	Tyr	Glu	Thr	Glu	Ala	Arg	Lys	Val	Phe	Cys	Glu	Val	Tyr		
	GGA	CTT	TTC	TAT	GAG	ACT	GAA	GCC	CGG	AAA	GTT	TTT	TGT	GAG	GTG	TAC	9	12
			275		-			280	- 3		- 2.		285			-		
,	-					Asn											J	
5	ርርጥ	GTG	ככר	GጥC	даа	AAC	ርፐር	AAG	CGC	CAG	GAΨ	GCC	CGA	ATC	ልጥሮ	GTG	ρ	64
				260					265					2,70				
	Glu	Ala	Gly		Glu	Ile	Thr	Phe	-	Gln	Ser	Phe	Phe		Asp	Pro		
	GAG	GCT	GGA	ATT	GAG	ATT	ACT	TTC	CGC	CAG	AGT	TTC	TTC	TCÀ	GAT	CCA	8	16
								-	•									

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			****	050					140									
	ıιρ	200	264	ULY.	<b></b> u					1	- , -	1	- <b></b>		- 3	<del>-</del>		
						GGC											10	<del>.</del>
	<b>m</b>	C.T.C	OMC.	CCC	Cmc	GGC	mmæ	a cm	CITIC	GGC	תיא רי	CCM	ጥርር	יטעע	_ _ _ _ _	ACC		824
40				580					585					590				v
	Tyr	His	Ile	Gly	Arg	Asn	Gln	Phe		Phe	Val	Суѕ	Gln		Arg	Leu		
	TAC	CAC	ATT	GGG	AGG	AAC	CAG	TTT	CCT	TTC	GTC	TGC	CAG	GCC	CGC	CTC	1	776
							•											
	2	- 4			565					570			-		575	-		
35						Leu				•							-	_ •
	GGC	ጥርር	ጥሮል	ርጥር	GCT	TTA	GCጥ	GCጥ	GTC	TTC	CCC	ርጥር	GGG	ርፓር	GAT	GGT	1'	728
	545					550					555					560		
	_	_	Ile	Gln	Asn	Ser		Pro	Asn	Leu		Asn	Leu	Thr	Ala			
30						TCA											16	680
		530				-,-	535					540				<b>-</b>		
						Cys											1	
25	ርጥር	רתיא	CCT	டேமா	GTC	TGT	ርጥር	ጥርር	արդուր	770	ልጥሮ	ጥልቦ	220	ፈጋጥ	САТ	СТС	1 4	532
			515					520					525					
	Ser	Gln	_	Leu	Phe	Ile	Ser		Ser	Val	Leu	Ser		Leu	Gly	Ile		
	TCA	CAG	AAA	CTC	TTT	ATC	TCC	GTC	TCA	GTT	CTC	TCC	AGC	CTG	GGC	ATT	15	584
20	~			500			<b></b>		505			<b></b>		510				
						Gln									_		1.	
	ጥርር	ccc	CCA	ርር ጥ	GAC	CAG	ACC	ርጥር	GTC	Δጥ⊂	ልልሮ	ልሮአ	الىلىل مىلىل	רפר	ጥጥር	ርጥር	1 '	536
					485					490					495			
15	Thr	Lys	Asp	Asp		Ser	Trp	Ser	Lys		Asp	Lys	Trp	Ile	_	Gly		
	ACC	AAG	GAT	GAT	CTT	TCC	TGG	TCC	AAA	ACA	GAT	AAA	TGG	ATT	GGA	GGG	14	188
	465	<b>G111</b>	neu	GIII	GIY	470	Jet	- Y -	LYS	n y o	475	GIY	* A *	+ Y +	nap	480		
10						GGC											14	140
10	C3.C	03.0	cam.	CNC	CCM	GGC	NCC.	ma c	220	አአሮ	אחשא	ccc	ma c	መአመ	CAC	300	1.	140
		450					455					460						
	His	Val	Val	Phe	Asp	Ala	Ser	Gly	Ser	Arg	Met	Ala	Trp	Thr	Leu	Ile		
	CAT	GTG	GTG	TTT	GAT	GCC	AGC	GGC	TCT	CGG	ATG	GCA	TGG	ACG	CTT	ATC	13	392
5																		
	<b>U111</b>	-76	435	9				440					445			~~1		
						ATG Met					_	_	_	_		_	1.	344
	•	<b>.</b>	m> 0	000	CC3	አመር	330	mcm.	mcc.	тсс	mmm.	C3.C	CCM	CMC	mom			

	III	265 1	WO	SEC					150								
	TCC	TCC	TAT	ATC	ACT	CTT	GTT	GTG	CTC	TTT	GTG	ccc	AAG	ATG	CGC	AGG	2352
			, , ,											•			
40	ser	GIU	755		MIG	AId	FIIE	760		n1a	261	Lea	765				•
40							TTT									Phe	2504
		<b></b>	a. c	<b>~</b>		000	(III)	000	mmm	666	mom	CMM	000	አመኦ	Cum	mm∕-	2304
				740					745					750			
							Ile										
35	GTG	GCA	GTC	CTG	TGC	CTC	ATC	ACT	GCT	CCT	GTC	ACC	ATG	ATT	CTG	TCC	2256
					123					,30					, , ,		
	Thr	Glu	Lys	ile	Asn 725	Asp	His	Arg	Ala	730	GIĀ	met	Alg	тте	735	ASI	
							CAC										2208
30															m: c		2222
	705					710					715					720	
	Leu	Leu	Leu	Leu	Gly	Ile	Phe	Leu	Ala	Tyr	Glu	Thr	Lys	Ser	Val	Ser	
	CTG	CTG	CTG	CTG	GGA	ATC	TTC	CTT	GCT	TAT	GAG	ACC	AAG	AGT	GTG	TCC	2160
23		550															
25	MIG	690	nec	van	1111	יבי	695	O-1	-10		~ } ~	700	-1.	-,, 5	1		
							Leu										2112
	`200	330	አሙን	מאג	7 C 7	тсс	CTT	ccc	ን ሙጥ	ጥጥር	ጥልጥ	උඋጥ	ጥልሮ	224	GGG	ርጥር	2112
			675					680					685				
20	Glu	Asp		Asp	Val	Ser	Ile		Pro	Gln	Leu	Glu		Суѕ	Ser	Ser	
							ATT										2064
		_ =		660					665			-		670			
							Ile										
15	GAC	ССТ	CTG	CAC	CGG	ACC	ATT	GAG	ACA	TTT	GCC	AAG	GAG	GAA	CCT	AAG	2016
					645					650					655		
	Leu	Leu	Val	Gly		Asp	Val	Leu	Thr		Ala	Ile	Trp	Gln		Val	
							GTC										1968
10																	
	625					630					635					640	
	Glu	Trp	Arg	Lys	Thr	Leu	Glu	Pro	Trp	Lys	Leu	Tyr	Ala	Thr	Val	Gly	
	GAG	TGG	AGG	AAG	ACT	CTG	GAA	ccc	TGG	AAG	CTG	TAT	GCC	ACA	GTG	GGC	1920
5		010					013					020					
	Lys	610	Trp	Trp	Val	HIS	Thr 615	vai	Pne	Thr	гуз	620	GIU	GIU	гàг	ьys	
							ACG										1872
	•		595					600				•	605	,,	·		******

	Ser	Ser 770	Tyr	Ile	Thr	Leu	Val 775	Val	Leu	Phe	Val	Pro 780	Lys	Met	Arg	Arg		-
	CTG	ATC	ACC	CGA	GGG	GAA	TGG	CAG	TCG	GAG	GCG	CAG	GAC	ACC	ATG	AAG	2	400
5	Leu	Ile	Thr	Arg	Gly	Glu	Trp	Gln	Ser	Glu	Ala	Gln	Asp	Thr	Met	Lys		
	785					790					795					800		
	ACA	GGG	TCA	TCG	ACC	AAC	AAC	AAC	GAG	GAG	GAG	AAG	TCC	CGG	CTG	TTG	2	448
	Thr	Gly	Ser	Ser	Thr	Asn	Asn	Asn	Glu	Glu	Glu	Lys	Ser	Arg	Leu	Leu		
10					805					810					815			
	GAG	AAG	GAG	AAC	CGT	GAA	CTG	GAA	AAG	ATC	ATT	GCT	GAG	AAA	GAG	GAG	2	496
	Glu	Lys	Glu	Asn	Arg	Glu	Leu	Glu	Lys	Ile	Ile	Ala	Glu	Lys	Glu	Glu		
				820					825					830				
15																		
	CGT	GTC	TCT	GAA	CTG	CGC	CAT	CAA	CTC	CAG	TCT	CGG	CAG	CAG	CTC	CGC	2	544
	Arg	Val	Ser	Glu	Leu	Arg	His	Gln	Leu	Gln	Ser	Arg	Gln	Gln	Leu	Arg		
			835					840					845					
20	TCC	CGG	CGC	CAC	CCA	CCG	ACA	ccc	CCA	GAA	CCC	TCT	GGG	GGC	CTG	CCC	2	592
	Ser	Arg	Arg	His	Pro	Pro	Thr	Pro	Pro	Glu	Pro	Ser	Gly	Gly	Leu	Pro		
		850					855					860						
	AGG	GGA	CCC	ССТ	GAG	ccc	ccc	GAC	CGG	СТТ	AGC	TGT	GAT	GGG	AGT	CGA	2	640
25	Arg	Gly	Pro	Pro	Glu	Pro	Pro	Asp	Arg	Leu	Ser	Cys	Asp	Gly	Ser	Arg		
	865					870					875					880		
	GTG	CAT	TTG	СТТ	TAT	AAG	TGA										2	661
	Val	His	Leu	Leu	Tyr	Lys	*											
30					885													
,																		

(2) INFORMATION FOR SEQ ID NO: 77:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 886 amino acids

(B) TYPE: amino acid(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 77:

35

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys Val Asn Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe Pro Ala Val Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile Leu Met Pro Gly Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser Ala Thr Leu His Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly Trp Lys Lys Ile Ala Thr Ile Gln Gln Thr

Thr Glu Val Phe Thr Ser Thr Leu Asp Asp Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile Thr Phe Arg Gln Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr 15 Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Ile Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Phe Glu Gly Val Ser Gly 

His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile

	Glu 465	Gln	Leu	Gln	Gly	Gly 470	Ser	Tyr	Lys	Lys	Ile 475	Gly	Tyr	Tyr	Asp	Ser 480
		_					_	_	_	_1	_	_			_,	
5	Thr	Lys	Asp	Asp	485	ser	Trp	Ser	гуs	490	Asp	rys	Trp	IIe	Gly 495	GΙΆ
	Ser	Pro	Pro	Ala 500	Asp	Gln	Thr	Leu	Val 505	Ile	Lys	Thr	Phe	Arg 510	Phe	Leu
10	Ser	Gln	Lys 515	Leu	Phe	Ile	Ser	Val 520	Ser	Val	Leu	Ser	Ser 525	Leu	Gly	Ile
15	Val	Leu 530	Ala	Val	Val	Cys	Leu 535	Ser	Phe	Asn	Ile	Туг 540	Asn	Ser	His	Val
	Arg 545	Tyr	Ile	Gln	Asn	Ser 550	Gln	Pro	Asn	Leu	Asn 555	Asn	Leu	Thr	Ala	Val 560
20	Gly	Cys	Ser	Leu	Ala 565	Leu	Ala	Ala	Val	Phe 570	Pro	Leu	Gly	Leu	Asp 575	Gly
25	Tyr	His	Ile	Gly 580	Arg	Asn	Gln	Phe	Pro 585	Phe	Val	Cys	Gln	Ala 590	Arg	Leu
	Trp	Leu	Leu 595	Gly	Leu	Gly	Phe	Ser 600	Leu	Gly	Туг	Gly	Ser 605	Met	Phe	Thr
30	Lys	Ile 610	Trp	Trp	Val	His	Thr 615	Val	Phe	Thr	Lys	Lys 620	Glu	Glu	Lys	Lys
	Glu 625	Trp	Arg	Lys	Thr	Leu 630	Glu	Pro	Trp	_	Leu 635	-	Ala	Thr	Val	Gly 6 <b>4</b> 0
35	Leu	Leu	Val	Gly	Met 645	Asp	Val	Leu	Thr	Leu 650	Ala	Ile	Trp	Gln	Ile 655	Val
40	Asp	Pro	Leu	His 660	Arg	Thr	Ile	G1u	Thr 665	Phe	Ala	Lys	Glu	Glu 670	Pro	Lys
· <del>·</del>	Glu	Asp	Ile 675	Asp	Val	Ser	Ile	Leu 680	Pro	Gln	Leu	Glu	His 685	Cys,	Ser	Ser

Arg Lys Met Asn Thr Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Gly Ile Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu Val Val Leu Phe Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu Ala Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu Arg His Gln Leu Gln Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro Pro Thr Pro Pro Glu Pro Ser Gly Gly Leu Pro Arg Gly Pro Pro Glu Pro Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr Lys 

(2) INFORMATION FOR SEQ ID NO: 78:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1692 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 5 (ii) MOLECULE TYPE: cDNA to mRNA (iii) HYPOTHETICAL: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1..1692 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 78: ATG TTG CTG CTG CTA CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC 48 Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly 5 10 GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC 96 Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile 25 ATA CAC CCG CCC TGG GAA GGG GGC ATC AGG TAC CGG GGC CTG ACT CGG 144 Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg 35 40 GAC CAG GTG AAG GCT ATC AAC TTC CTG CCA GTG GAC TAT GAG ATT GAG 192 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 55 60 TAT GTG TGC CGG GGG GAG CGC GAG GTG GTG GGG CCC AAG GTC CGC AAG 240 Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys TGC CTG GCC AAC GGC TCC TGG ACA GAT ATG GAC ACA CCC AGC CGC TGT 288 Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys 85 GTC CGA ATC TGC TCC AAG TCT TAT TTG ACC ATT ACC GAC CAA ATC TAC 336 H1865-1 WO SEQ 156

	Val	Arg	Ile	Cys 100	Ser	Lys	Ser	Tyr	Leu 105	Thr	Ile	Thr	Asp	Gln 110	Ile	Туř	
	CGG	GCA	ATG	AAC	TCT	TCG	TCC	TTT	GAG	GGT	GTC	TCT	GGC	CAT	GTG	GTG	384
5	Arg	Ala	Met	Asn	Ser	Ser	Ser	Phe	Glu	Gly	Val	Ser	Gly	His	Val	Val	
			115					120					125				
	ттт	GAT	GCC	AGC	GGC	TCT	CGG	ATG	GCA	TGG	ACG	CTT	ATC	GAG	CAG	CTT	432
	Phe	Asp	Ala	Ser	Gly	Ser	Arg	Met	Ala	Trp	Thr	Leu	Ile	Glu	Gln	Leu	
10		130					135					140					
	CAG	GGT	GGC	AGC	TAC	AAG	AAG	ATT	GGC	TAC	TAT	GAC	AGC	ACC	AAG	GAT	480
	Gln	Gly	Gly	Ser	Tyr	_	Lys	Ile	Gly	Tyr	Tyr	Asp	Ser	Thr	Lys	_	
	145					150					155					160	
15																	500
	_												GGG				528
	Asp	Leu	ser	Trp	165	ьўs	THE	Asp	гуѕ	170	TTE	GIĀ	Gly	Ser		Pro	
					103					170					175		
20	CCT	GAC	CAG	ACC.	СТС	ርጥር	ልጥር	AAG	ACA	ጥጥ	CGC	שיזיי	CTG	ጥርል	CAG	ΔΔΑ	576
20						_	_						Leu		_		• • •
				180				-4-	185		5			190			
	CTC	ттт	ATC	TCC	GTC	TCA	GTT	CTC	TCC	AGC	CTG	GGC	ATT	GTC	CTA	GCT	624
25	Leu	Phe	Ile	Ser	Val	Ser	Val	Leu	Ser	Ser	Leu	Gly	Ile	Val	Leu	Ala	
			195					200					205				
	GTT	GTC	TGT	CTG	TCC	TTT	AAC	ATC	TAC	AAC	TCA	CAT	GTC	CGT	TAT	ATC	672
	Val	Val	Cys	Leu	Ser	Phe	Asn	Ile	Tyr	Asn	Ser	His	Val	Arg	Tyr	Ile	
30		210					215					220					
													GTG				720
	-	Asn	Ser	Gln	Pro		Leu	Asn	Asn	Leu		Ala	Val	Gly	Суз		
	225					230					235					240	
35						CMC	mma	000	ama	000	oma	~ N M		ma.c	CAC	አ ሙጠ	768
													GGT				700
	Leu	Ala	Leu	AIG		vai	rne	PLO	rea	250	nea	ASD	Gly	IAT	255	116	
					245					430					ددے		
40	GGG	AGG	ልልሮ	CAG	ጥጥጥ	CCT	ፐጥ	GTC	TGC	CAG	GCC	ርርር	CTC	TGG	CTC	CTG	816,
40													Leu				U
	J_Y	9		260					265			9		270			

													•				
	ĠGC	CTG	GGC	TTT	AGT	CTG	GGC	TAC	GGT	TCC	ATG	TTC	ACC	AAG	ATT	TGG	864
	Gly	Leu	Gly	Phe	Ser	Leu	Gly	Tyr	Gly	Ser	Met	Phe	Thr	Lys	Ile	Trp	
			275					280					285				
5	TGG	GTC	CAC	ACG	GTC	TTC	ACA	AAG	AAG	GAA	GAA	AAG	AAG	GAG	TGG	AGG	912
	Trp	Val	His	Thr	Val	Phe	Thr	Lys	Lys	Glu	Glu	Lys	Lys	Glu	Trp	Arg	
		290					295					300					
	AAG	ACT	CTG	GAA	ccc	TGG	AAG	CTG	TAT	GCC	ACA	GTG	GGC	CTG	CTG	GTG	960
10	Lys	Thr	Leu	Glu	Pro	Trp	Lys	Leu	Tyr	Ala	Thr	Val	Gly	Leu	Leu	Val	
	305					310					315					320	
	GGC	ATG	GAT	GTC	CTC	ACT	CTC	GCC	ATC	TGG	CAG	ATC	GTG	GAC	CCT	CTG	1008
	Gly	Met	Asp	Va1	Leu	Thr	Leu	Ala	Ile	Trp	Gln	Ile	Val	Asp	Pro	Leu	
15					325					330					335		
	CAC	CGG	ACC	ATT	GAG	ACA	TTT	GCC	AAG	GAG	GAA	CCT	AAG	GAA	GAT	ATT	1056
	His	Arg	Thr	Ile	Glu	Thr	Phe	Ala	Lys	Glu	Glu	Pro	Lys	Glu	Asp	Ile	
				340					345					350			
20																	
	GAC	GTC	TCT	ATT	CTG	CCC	CAG	CTG	GAG	CAT	TGC	AGC	TCC	AGG	AAG	ATG	1104
	Asp	Val	Ser	Ile	Leu	Pro	Gln	Leu	Glu	His	Суѕ	Ser	Ser	Arg	Lys	Met	
			355					360					365				
25	AAT	ACA	TGG	CTT	GGC	ATT	TTC	TAT	GGT	TAC	AAG	GGG	CTG	CTG	CTG	CTG	1152
	Asn	Thr	Trp	Leu	Gly	Ile	Phe	Tyr	Gly	Tyr	Lys	Gly	Leu	Leu	Leu	Leu	
		370					375					380					
												GTG					1200
30	Leu	Gly	Ile	Phe	Leu	Ala	Tyr	Glu	Thr	Lys		Val	Ser	Thr	Glu		
	385					390					395					400	
																	4040
												TAC					1248
	Ile	Asn	Asp	His	Arg	Ala	Val	Gly	Met		Ile	Tyr	Asn	Val		Val	
35					405					410					415		
															222	<b>63.6</b>	1296
																CAG	1230
	Leu	Cys	Leu			Ala	Pro	val			TIE	Leu	ser		GIN	Gln	
				420					425					430			
40			_	_					-	000			mma	maa	mcc.	መጽሞ	1344
																TAT	7744
	Asp				Ala	Phe	Ala			ATA	тте	val			ser	Tyr	
		•	435					440					445				

	•																-—
	ATC	ACT	CTT	GTT	GTG	CTC	TTT	GTG	CCC	AAG	ATG	CGC	AGG	CTG	ATC	ACC	1392
	Ile	Thr	Leu	Val	Val	Leu	Phe	Val	Pro	Lys	Met	Arg	Arg	Leu	Ile	Thr	
		450					455					460					
5																	
	CGA	GGG	GAA	TGG	CAG	TCG	GAG	GCG	CAG	GAC	ACC	ATG	AAG	ACA	GGG	TCA	1440
	Arg	Gly	Glu	Trp	Gln	Ser	Glu	Ala	Gln	Asp	Thr	Met	Lys	Thr	Gly	Ser	
	465					470					475					480	
10	TCG	ACC	AAC	AAC	AAC	GAG	GAG	GAG	AAG	TCC	CGG	CTG	TTG	GAG	AAG	GAG	1488
	Ser	Thr	Asn	Asn	Asn	G1u	Glu	Glu	Lys	Ser	Arg	Leu	Leu	Glu	Lys	Glu	
					485					490					495		
	AAC	CGT	GAA	CTG	GAA	AAG	ATC	ATT	GCT	GAG	AAA	GAG	GAG	CGT	GTC	TCT	1536
15	Asn	Arg	Glu	Leu	Glu	Lys	Ile	Ile	Ala	Glu	Lys	Glu	Glu	Arg	Val	Ser	
				500					505					510			
	GAA	CTG	CGC	CAT	CAA	CTC	CAG	TCT	CGG	CAG	CAG	CTC	CGC	TCC	CGG	CGC	1584
	Glu	Leu	Arg	His	Gln	Leu	Gln	Ser	Arg	Gln	Gln	Leu	Arg	Ser	Arg	Arg	:
20			515					520					525				
	CAC	CCA	CCG	ACA	ccc	CCA	GAA	CCC	TCT	GGG	GGC	CTG	ccc	AGG	GGA	CCC	1632
	His	Pro	Pro	Thr	Pro	Pro	Glu	Pro	Ser	Gly	Gly	Leu	Pro	Arg	Gly	Pro	
		530					535					540					
25																	
	CCT	GAG	ccc	CCC	GAC	CGG	CTT	AGC	TGT	GAT	GGG	AGT	CGA	GTG	CAT	TTG	1680
	Pro	Glu	Pro	Pro	Asp	Arg	Leu	Ser	Суз	Asp	Gly	Ser	Arg	Val	His	Leu	
	545					550					555					560	
30	CTT	TAT	AAG	TGA													1692
	Leu	Tyr	Lys	*													
		-	_														

## (2) INFORMATION FOR SEQ ID NO: 79:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 563 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 79:

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly
1 5 10 15

5

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile
20 25 30

...

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg
35 40 45

10

Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu
50 55 60

**т** 

Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys

6

Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys
85 90 95

20

Val Arg Ile Cys Ser Lys Ser Tyr Leu Thr Ile Thr Asp Gln Ile Tyr
100 105 110

25

Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val
115 120 125

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Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu 130 135 140

30

Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp 145 150 155 160

Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro 165 170 175

35

Ala Asp Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys 180 185 190

Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala 195 200 205

40

Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile 210 215 220

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Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Asn Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Arg Lys Met Asn Thr Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Leu Gly Ile Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr 

Ile Thr Leu Val Val Leu Phe Val Pro Lys Met Arg Arg Leu Ile Thr 455 460 Arg Gly Glu Trp Gln Ser Glu Ala Gln Asp Thr Met Lys Thr Gly Ser 470 475 465 Ser Thr Asn Asn Asn Glu Glu Glu Lys Ser Arg Leu Leu Glu Lys Glu 485 490 495 Asn Arg Glu Leu Glu Lys Ile Ile Ala Glu Lys Glu Glu Arg Val Ser 500 505 510 Glu Leu Arg His Gln Leu Gln Ser Arg Gln Gln Leu Arg Ser Arg Arg 515 520 15 His Pro Pro Thr Pro Pro Glu Pro Ser Gly Gly Leu Pro Arg Gly Pro 535 540 530 Pro Glu Pro Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His Leu 560 550 555 20 545 Leu Tyr Lys 25 (2) INFORMATION FOR SEQ ID NO: 80: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2602 base pairs (B) TYPE: nucleic acid 30 (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: cDNA to mRNA 35 (iii) HYPOTHETICAL: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens 40 (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1...318

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 80:

	ATG	TTG	CTG	CTG	CTG	CTA	CTG	GCG	CCA	CTC	TTC	CTC	CGC	CCC	CCG	GGC	48
5	Met	Leu	Leu	Leu	Leu	Leu	Leu	Ala	Pro	Leu	Phe	Leu	Arg	Pro	Pro	Gly	
	1				5					10					15		
															CAG		96
	Ala	Gly	Gly	Ala	Gln	Thr	Pro	Asn		Thr	Ser	Glu	Gly		Gln	Ile	
10				20					25					30			
									. ma	100	m. a		000	OMC.	» cm	000	144
															ACT Thr		144
	TTE	HIS	35	Pro	Trp	GIU	GIY	40	TIE	Arg	ıyı	ALG	45	neu	1111	nrg	
15			33					40					43				
13	GAC	CAG	GTG	AAG	GCT	ATC	AAC	TTC	CTG	CCA	GTG	GAC	TAT	GAG	ATT	GAT	192
															Ile		
	<b>-</b>	50		-			55					60					
20	GAA	TCG	AAC	GCC	ACA	СТС	AGA	ACG	GCG	CGC	AGT	GTA	CAT	CGG	GGC	ACT	240
	Glu	Ser	Asn	Ala	Thr	Leu	Arg	Thr	Ala	Arg	Ser	Val	His	Arg	Gly	Thr	
	65					70					75					80	
															GCC		288
25	Val	Ser	His	Glu		Gly	Leu	Ala	Arg		Pro	Gly	Leu	Pro	Ala	Arg	
					85					90					95		
		221	0 h m	~~~	00m	00 N	CCA	CCM	C 3 3	ma C	ccc	~» ~~	720	אתיייי	TGCC	30	338
						Gly				±	CCG	CAGG	JAC A	HICC	1GCC	33	330
30	GIĀ	GIY	ASP	100	Ala	GIY	GIŞ	ALG	105								
30				100													
	ACT.	ATGA	GCT	CAAG	CTCA'	TC C.	ACCA	CGAC	A GC	AAGT	GTGA	TCC	AGGC	CAA	GCCA	CCAAGT	398
	ACC	ТАТА	TGA	GCTG	CTCT.	AC A	ACGA	CCCT	A TC	AAGA'	TCAT	CCT	TATG	CCT	GGCT	GCAGCT	458
35																	
	CTG	TCTC	CAC	GCTG	GTGG	CT G	AGGC'	TGCT.	A GG	ATGT	GGAA	CCT	CATT	GTG	CTTT	CCTATG	518
	GCT	CCAG	CTC	ACCA	GCCC	TG T	CAAA	CCGG	C AG	CGTT'	TCCC	CAC	TTTC	TTC	CGAA	CGCACÇ	578
40	CAT	CAGC	CAC	ACTC	CACA	AC C	CTAC	CCGC	G TG	AAAC	TCTT	TGA	AAAG	TGG	GGCT	GGAAGA	638 <sub>v</sub>
								_,						<b></b> -	<b>.</b>	maa: ==	600
	AGA	TTGC	TAC	CATC	CAGC	AG A	CCAC	TGAG	G TC	TTÇA	CTTC	GAC	TCTG	GAC	GACC	TGGAGG	698

	AACGAGTGAA	GGAGGCTGGA	ATTGAGATTA	CTTTCCGCCA	GAGTTTCTTC	TCAGATCCAG	758
	CTGTGCCCGT	CAAAAACCTG	AAGCGCCAGG	ATGCCCGAAT	CATCGTGGGA	CTTTTCTATG	818
5	AGACTGAAGC	CCGGAAAGTT	TTTTGTGAGG	TGTACAAGGA	GCGTCTCTTT	GGGAAGAAGT	878
	ACGTCTGGTT	CCTCATTGGG	TGGTATGCTG	ACAATTGGTT	CAAGATCTAC	GACCCTTCTA	. 938
10	TCAACTGCAC	AGTGGATGAG	ATGACTGAGG	CGGTGGAGGG	CCACATCACA	ACTGAGATTG	998
	TCATGCTGAA	TCCTGCCAAT	ACCCGCAGCA	TTTCCAACAT	GACATCCCAG	GAATTTGTGG	1058
	AGAAACTAAC	CAAGCGACTG	AAAAGACACC	CTGAGGAGAC	AGGAGGCTTC	CAGGAGGCAC	1118
15	CGCTGGCCTA	TGATGCCATC	TGGGCCTTGG	CACTGGCCCT	GAACAAGACA	TCTGGAGGAG	1178
	GCGGCCGTTC	TGGTGTGCGC	CTGGAGGACT	TCAACTACAA	CAACCAGACC	ATTACCGACC	1238
20	AAATCTACCG	GGCAATGAAC	TCTTCGTCCT	TTGAGGGTGT	CTCTGGCCAT	GTGGTGTTTG	1298
	ATGCCAGCGG	CTCTCGGATG	GCATGGACGC	TTATCGAGCA	GCTTCAGGGT	GGCAGCTACA	1358
	AGAAGATTGG	CTACTATGAC	AGCACCAAGG	ATGATCTTTC	CTGGTCCAAA	ACAGATAAAT	1418
25	GGATTGGAGG	GTCCCCCCA	GCTGACCAGA	CCCTGGTCAT	CAAGACATTC	CGCTTCCTGT	1478
	CACAGAAACT	CTTTATCTCC	GTCTCAGTTC	TCTCCAGCCT	GGGCATTGTC	CTAGCTGTTG	1538
30	TCTGTCTGTC	CTTTAACATC	TACAACTCAC	ATGTCCGTTA	TATCCAGAAC	TCACAGCCCA	1598
	ACCTGAACAA	CCTGACTGCT	GTGGGCTGCT	CACTGGCTTT	AGCTGCTGTC	TTCCCCCTGG	1658
	GGCTCGATGG	TTACCACATT	GGGAGGAACC	AGTTTCCTTT	CGTCTGCCAG	GCCCGCCTCT	1718
35	GGCTCCTGGG	CCTGGGCTTT	AGTCTGGGCT	ACGGTTCCAT	GTTCACCAAG	ATTTGGTGGG	1778
	TCCACACGGT	CTTCACAAAG	AAGGAAGAAA	AGAAGGAGTG	GAGGAAGACT	CTGGAACCCT	1838
40	GGAAGCTGTA	TGCCACAGTG	GGCCTGCTGG	TGGGCATGGA	TGTCCTCACT	CTCGCCATCT	1898
	GGCAGATCGT	GGACCCTCTG	CACCGGACCA	TTGAGACATT	TGCCAAGGAG	GAACCTAAGG	1958
	AAGATATTGA	CGTCTCTATT	CTGCCCCAGC	TGGAGCATTG	CAGCTCCAGG	AAGATGAATA	2018

	CATGGCTTGG	CATTTTCTAT	GGTTACAAGG	GGCTGCTGCT	GCTGCTGGGA	ATCTTCCTTG	2078
5	CTTATGAGAC	CAAGAGTGTG	TCCACTGAGA	AGATCAATGA	TCACCGGGCT	GTGGGCATGG	2138
J	СТАТСТАСАА	TGTGGCAGTC	CTGTGCCTCA	TCACTGCTCC	TGTCACCATG	ATTCTGTCCA	2198
	GCCAGCAGGA	TGCAGCCTTT	GCCTTTGCCT	CTCTTGCCAT	AGTTTTCTCC	TCCTATATCA	2258
0	CTCTTGTTGT	GCTCTTTGTG	CCCAAGATGC	GCAGGCTGAT	CACCCGAGGG	GAATGGCAGT	2318
	CGGAGGCGCA	GGACACCATG	AAGACAGGGT	CATCGACCAA	CAACAACGAG	GAGGAGAAGT	2378
5	CCCGGCTGTT	GGAGAAGGAG	AACCGTGAAC	TGGAAAAGAT	CATTGCTGAG	AAAGAGGAGC	2438
J	GTGTCTCTGA	ACTGCGCCAT	CAACTCCAGT	CTCGGCAGCA	GCTCCGCTCC	CGGCGCCACC	2498
	CACCGACACC	CCCAGAACCC	TCTGGGGGCC	TGCCCAGGGG	ACCCCCTGAG	CCCCCGACC	2558
0	GGCTTAGCTG	TGATGGGAGT	CGAGTGCATT	TGCTTTATAA	GTGA		2602

#### (2) INFORMATION FOR SEQ ID NO: 81:

25 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 105 amino acids

(B) TYPE: amino acid(D) TOPOLOGY: linear

- 30 (ii) MOLECULE TYPE: protein
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 81:

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly
35 1 5 10 15

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile
20 25 30

40 Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg
35 40 45

Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Asp

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50 55 60

Glu Ser Asn Ala Thr Leu Arg Thr Ala Arg Ser Val His Arg Gly Thr
65 70 75 80

Val Ser His Glu Arg Gly Leu Ala Arg Gly Pro Gly Leu Pro Ala Arg 85 90 95

Gly Gly Asp Gly Ala Gly Gly Arg Glu
10 100 105

- (2) INFORMATION FOR SEQ ID NO: 82:
- 15 (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2367 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA to mRNA

- (iii) HYPOTHETICAL: NO
- 25 (vi) ORIGINAL SOURCE:

20

- (A) ORGANISM: Homo sapiens
- (ix) FEATURE:
  - (A) NAME/KEY: CDS
- 30 (B) LOCATION:1..294
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 82:

ATG TTG CTG CTG CTG CTA CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC

48

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly

1 5 10 15

GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile

20

25

30

ATA CAC CCG CCC TGG GAA GGG GGC ATC AGG TAC CGG GGC CTG ACT CGG 144

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg

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	. 3	5 .	40		45	• •••	
				CTG CCA GTG Leu Pro Val			19
5	50		55		60		
		s Arg Gly G	lu Arg Glu	GTG GTG GGG Val Val Gly			. 24
10	65		70	75		80	
				GAT ATG GAC Asp Met Asp 90			28
15	GTG TGA TC		CACCAAGT AC	CTATATGA GCT	rgctctac aac		34
20	TCAAGATCAT	CCTTATGCCT	GGCTGCAGCT	CTGTCTCCAC	GCTGGTGGCT	GAGGCTGCTA	40
20	GGATGTGGAA	CCTCATTGTG	CTTTCCTATG	GCTCCAGCTC	ACCAGCCCTG	TCAAACCGGC	• 46
	AGCGTTTCCC	CACTTTCTTC	CGAACGCACC	CATCAGCCAC	ACTCCACAAC	CCTACCCGCG	52
25	TGAAACTCTT	TGAAAAGTGG	GGCTGGAAGA	AGATTGCTAC	CATCCAGCAG	ACCACTGAGG	58
	TCTTCACTTC	GACTCTGGAC	GACCTGGAGG	AACGAGTGAA	GGAGGCTGGA	ATTGAGATTA	64
30	CTTTCCGCCA	GAGTTTCTTC	TCAGATCCAG	CTGTGCCCGT	CAAAAACCTG	AAGCGCCAGG	70
	ATGCCCGAAT	CATCGTGGGA	CTTTTCTATG	AGACTGAAGC	CCGGAAAGTT	TTTTGTGAGG	76
	TGTĄCAAGGA	GCGTCTCTTT	GGGAAGAAGT	ACGTCTGGTT	CCTCATTGGG	TGGTATGCTG	82
35	ACAATTGGTT	CAAGATCTAC	GACCCTTCTA	TCAACTGCAC	AGTGGATGAG	ATGACTGAGG	88
	CGGTGGAGGG	CCACATCACA	ACTGAGATTG	TCATGCTGAA	TCCTGCCAAT	ACCCGCAGCA	94
40	TTTCCAACAT	GACATCCCAG	GAATTTGTGG	AGAAACTAAC	CAAGCGACTG	AAAAGACACC	100
	CTGAGGAGAC	AGGAGGCTTC	CAGGAGGCAC	CGCTGGCCTA	TGATGCCATC	TGGGCCTTGG	106
	CACTGGCCCT	GAACAAGACA	TCTGGAGGAG	GCGGCCGTTC	TGGTGTGCGC	CTGGAGGACT	112

	TCAACTACAA	CAACCAGACC	ATTACCGACC	AAATCTACCG	GGCAATGAAC	TCTTCGTCCT	118
5	TTGAGGGTGT	CTCTGGCCAT	GTGGTGTTTG	ATGCCAGCGG	CTCTCGGATG	GCATGGACGC	124
3	TTATCGAGCA	GCTTCAGGGT	GGCAGCTACA	AGAAGATTGG	CTACTATGAC	AGCACCAAGG	1304
	ATGATCTTTC	CTGGTCCAAA	ACAGATAAAT	GGATTGTTAT	ATCCAGAACT	CACAGCCCAA	1364
0	CCTGAACAAC	CTGACTGCTG	TGGGCTGCTC	ACTGGCTTTA	GCTGCTGTCT	TCCCCTGGG	142
	GCTCGATGGT	TACCACATTG	GGAGGAACCA	GTTTCCTTTC	GTCTGCCAGG	CCCGCCTCTG	1484
.5	GCTCCTGGGC	CTGGGCTTTA	GTCTGGGCTA	CGGTTCCATG	TTCACCAAGA	TTTGGTGGGT	154
. •	CCACACGGTC	TTCACAAAGA	AGGAAGAAAA	GAAGGAGTGG	AGGAAGACTC	TGGAACCCTG	1604
	GAAGCTGTAT	GCCACAGTGG	GCCTGCTGGT	GGGCATGGAT	GTCCTCACTC	TCGCCATCTG	1664
.0	GCAGATCGTG	GACCCTCTGC	ACCGGACCAT	TGAGACATTT	GCCAAGGAGG	AACCTAAGGA	1724
	AGATATTGAC	GTCTCTATTC	TGCCCCAGCT	GGAGCATTGC	AGCTCCAGGA	AGATGAATAC	178
25	ATGGCTTGGC	ATTTTCTATG	GTTACAAGGG	GCTGCTGCTG	CTGCTGGGAA	TCTTCCTTGC	1844
•	TTATGAGACC	AAGAGTGTGT	CCACTGAGAA	GATCAATGAT	CACCGGGCTG	TGGGCATGGC	1904
	TATCTACAAT	GTGGCAGTCC	TGTGCCTCAT	CACTGCTCCT	GTCACCATGA	TTCTGTCCAG	1964
10	CCAGCAGGAT	GCAGCCTTTG	CCTTTGCCTC	TCTTGCCATA	GTTTTCTCCT	CCTATATCAC	2024
	TCTTGTTGTG	CTCTTTGTGC	CCAAGATGCG	CAGGCTGATC	ACCCGAGGGG	AATGGCAGTC	2084
35	GGAGGCGCAG	GACACCATGA	AGACAGGGTC	ATCGACCAAC	AACAACGAGG	AGGAGAAGTC	2144
,5	CCGGCTGTTG	GAGAAGGAGA	ACCGTGAACT	GGAAAAGATC	ATTGCTGAGA	AAGAGGAGCG	2204
	TGTCTCTGAA	CTGCGCCATC	AACTCCAGTC	TCGGCAGCAG	CTCCGCTCCC	GGCGCCACCC	2264
<b>\$</b> 0	ACCGACACCC	CCAGAACCCT	CTGGGGGCCT	GCCCAGGGGA	CCCCTGAGC	CCCCGACCG	2324
	GCTTAGCTGT	GATGGGAGTC	GAGTGCATTT	GCTTTATAAG	TGA	•	2367

PCT/SE98/01947 WO 99/21890

(2) INFORMATION FOR SEQ ID NO: 83:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 97 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

10

15

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 83:

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly 1

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile 25 20

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg 35 40 45

Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 55 60

Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys 70 75 65

Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys 85 90

30

40

Val

- (2) INFORMATION FOR SEQ ID NO: 84:
  - (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2489 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA to mRNA

H1865-1 WO SEQ

(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO 5 (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (ix) FEATURE: (A) NAME/KEY: CDS 10 (B) LOCATION: 1..1491 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 84: 15 ATG TTG CTG CTG CTA CTG GCG CCA CTC TTC CTC CGC CCC CCG GGC 48 Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly 1 5 10 15 GCG GGC GGG GCG CAG ACC CCC AAC GCC ACC TCA GAA GGT TGC CAG ATC 96 Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile ATA CAC CCG CCC TGG GAA GGG GGC ATC AGG TAC CGG GGC CTG ACT CGG 144 Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg 35 GAC CAG GTG AAG GCT ATC AAC TTC CTG CCA GTG GAC TAT GAG ATT GAG 192 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu 50 55 60 30 TAT GTG TGC CGG GGG GAG CGC GAG GTG GGG CCC AAG GTC CGC AAG 240 Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys 65 70 75 TGC CTG GCC AAC GGC TCC TGG ACA GAT ATG GAC ACA CCC AGC CGC TGT 288 Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys 85 90 GTC CGA ATC TGC TCC AAG TCT TAT TTG ACC CTG GAA AAT GGG AAG GTT 336 Val Arg Ile Cys Ser Lys Ser Tyr Leu Thr Leu Glu Asn Gly Lys Val 100 105 384 TTC CTG ACG GGT GGG GAC CTC CCA GCT CTG GAC GGA GCC CGG GTG GAT H1865-1 WO SEQ 170

	Phe	Leu	Thr 115	Gly	Gly	Asp	Leu	Pro 120	Ala	Leu	Asp	Glý	Ala 125	Arg	Val	Asp		
	TTC	CGG	TGT	GAC	ccc	GAC	TTC	CAT	CTG	TGT	GAT	CCA	GGC	CAA	GCC	ACC		432
5	Phe	Arg 130	Суѕ	Asp	Pro	Asp	Phe 135	His	Leu	Cys	Asp	Pro 140	Gly	Gln	Ala	Thr		
	AAG	TAC	CTA	тат	GAG	CTG	CTC	TAC	AAC	GAC	ССТ	ATC	AAG	ATC	ATC	CTT	•	480
	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro	Ile	Lys	Ile	Ile	Leu		
10	145					150					155					160		
	ATG	CCT	GGC	TGC	AGC	TCT	GTC	TCC	ACG	CTG	GTG	GCT	GAG	GCT	GCT	AGG		528
	Met	Pro	Gly	Суѕ	Ser	Ser	Val	Ser	Thr	Leu	Val	Ala	Glu	Ala	Ala	Arg		
15					165					170					175			
13	ATG	TGG	AAC	CTC	АТТ	GTG	CTT	TCC	TAT	GGC	TCC	AGC	TCA	CCA	GCC	CTG		576
	Met	Trp	Asn	Leu	Ile	Val	Leu	Ser	Tyr	Gly	Ser	Ser	Ser	Pro	Ala	Leu		
				180					185					190				
20	TCA	AAC	CGG	CAG	CGT	TTC	CCC	ACT	TTC	TTC	CGA	ACG	CAC	CCA	TCA	GCC		624
	Ser	Asn	Arg	Gln	Arg	Phe	Pro	<b>Thr</b>	Phe	Phe	Arg	Thr	His	Pro	Ser	Ala		
			195					200					205				·	
	ACA	CTC	CAC	AAC	CCT	ACC	CGC	GTG	AAA	CTC	TTT	GAA	AAG	TGG	GGC	TGG		672
25	Thr	Leu	His	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe	Glu	Lys	Trp	Gly	Trp		
		210					215					220						
	AAG	AAG	ATT	GCT	ACC	ATC	CAG	CAG	ACC	ACT	GAG	GTC	TTC	ACT	TCG	ACT		720
	Lys	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	Val	Phe	Thr	Ser	Thr		
30	225					230					235					240		
	CTG	GAC	GAC	CTG	GAG	GAA	CGA	GTG	AAG	GAG	GCT	GGA	ATT	GAG	АТТ	ACT		768
	Leu	Asp	Asp	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala	Gly	Ile	Glu	Ile	Thr		
					245					250					255			
35	mmc	ccc	CAC	» Cm	mmc	TTC	መሮአ	CAM	CCA	CCT	CTC	ccc	CTC	222	7 7 C	CTC		816
						Phe												010
	1110	g	<b>0111</b>	260			502	1.0,0	265		<b>,</b>		vul	270		204		
40						CGA												864 <sub>v</sub>
	Lys	Arg		Asp	Ala	Arg	Ile		Val	GIA	Leu	Phe		Glu	Thr	GIu		
			275					280					285					

									•								
	GCC	CGG	AAA	GTT	TTT	TGT	GAG	GTG	TAC	AAG	GAG	CGT	CŤC	TTT	GGG	AAG	912
	Ala	Arg	Lys	Val	Phe	Cys	Glu	Val	Tyr	Lys	Glu	Arg	Leu	Phe	Gly	Lys	
		290					295					300					
5	AAG	TAC	GTC	TGG	TTC	CTC	ATT	GGG	TGG	TAT	GCT	GAC	AAT	TGG	TTC	AAG	960
	Lys	Tyr	Val	Trp	Phe	Leu	Ile	Gly	Trp	Tyr	Ala	Asp	Asn	Trp	Phe	Lys	
	305					310					315					320	
	ATC	TAC	GAC	CCT	TCT	ATC	AAC	TGC	ACA	GTG	GAT	GAG	ATG	ACT	GAG	GCG	1008
10	Ile	Tyr	Asp	Pro	Ser	Ile	Asn	Cys	Thr	Val	Asp	Glu	Met	Thr	Glu	Ala	
					325					330					335		
	GTG	GAG	GGC	CAC	ATC	ACA	ACT	GAG	ATT	GTC	ATG	CTG	AAT	CCT	GCC	AAT	1056
	Val	Glu	Gly	His	Ile	Thr	Thr	Glu	Ile	Val	Met	Leu	Asn	Pro	Ala	Asn	
15				340					345					350			
										-							
	ACC	CGC	AGC	ATT	TCC	AAC	ATG	ACA	TCC	CAG	GAA	TTT	GTG	GAG	AAA	CTA	1104
	Thr	Arg	Ser	Ile	Ser	Asn	Met	Thr	Ser	Gln	Glu	Phe	Val	Glu	Lys	Leu	
			355					360					365				
20																	
	ACC	AAG	CGA	CTG	AAA	AGA	CAC	CCT	GAG	GAG	ACA	GGA	GGC	TTC	CAG	GAG	1152
	Thr	Lys	Arg	Leu	Lys	Arg	His	Pro	Glu	Glu	Thr	Gly	Gly	Phe	Gln	Glu	•
		370					375					380					
25	GCA	CCG	CTG	GCC	TAT	GAT	GCC	ATC	TGG	GCC	TTG	GCA	CTG	GCC	CTG	AAC	1200
	Ala	Pro	Leu	Ala	Tyr	Asp	Ala	Ile	Trp	Ala	Leu	Ala	Leu	Ala	Leu	Asn	
	385					390					395					400	
	AAG	ACA	TCT	GGA	GGA	GGC	GGC	CGT	TCT	GGT	GTG	CGC	CTG	GAG	GAC	TTC	1248
30	Lys	Thr	Ser	Gly	Gly	Gly	Gly	Arg	Ser	Gly	Val	Arg	Leu	Glu	Asp	Phe	
					405					410					415		
	AAC	TAC	AAC	AAC	CAG	ACC	ATT	ACC	GAC	CAA	ATC	TAC	CGG	GCA	ATG	AAC	1296
	Asn	Tyr	Asn	Asn	Gln	Thr	Ile	Thr	Asp	Gln	Ile	Tyr	Arg	Ala	Met	Asn	
35				420					425					430			
	TCT	TCG	TCC	TTT	GAG	GGT	GTC	TCT	GGC	CAT	GTG	GTG	TTT	GAT	GCC	AGC	1344
	Ser	Ser	Ser	Phe	Glu	Gly	Val	Ser	Gly	His	Val	Val	Phe	Asp	Ala	Ser	
			435					440					445				
40																	υ
	GGC	TCT	CGG	ATG	GCA	TGG	ACG	CTT	ATC	GAG	CAG	CTT	CAG	GGT	GGÇ	AGC	1392
	Gly	Ser	Arg	Met	Ala	Trp	Thr	Leu	Ile	Glu	Gln	Leu	Gln	Gly	Gly	Ser	•
		450					455					460					

5	TAC AAG AAG ATT GGC TAC TAT GAC AGC ACC AAG GAT GAT CTT TCC TGG Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp 465 470 475 480	1440
	TCC AAA ACA GAT AAA TGG ATT GTT ATA TCC AGA ACT CAC AGC CCA ACC Ser Lys Thr Asp Lys Trp Ile Val Ile Ser Arg Thr His Ser Pro Thr 485 490 495	1488
10	TGA ACAACCTGAC TGCTGTGGGC TGCTCACTGG CTTTAGCTGC TGTCTTCCCC	1541
15	CTGGGGCTCG ATGGTTACCA CATTGGGAGG AACCAGTTTC CTTTCGTCTG CCAGGCCCGC	1601
	CTCTGGCTCC TGGGCCTGGG CTTTAGTCTG GGCTACGGTT CCATGTTCAC CAAGATTTGG	1661
	TGGGTCCACA CGGTCTTCAC AAAGAAGGAA GAAAAGAAGG AGTGGAGGAA GACTCTGGAA	1721
20	CCCTGGAAGC TGTATGCCAC AGTGGGCCTG CTGGTGGGCA TGGATGTCCT CACTCTCGCC	1781
	ATCTGGCAGA TCGTGGACCC TCTGCACCGG ACCATTGAGA CATTTGCCAA GGAGGAACCT	1841
25	AAGGAAGATA TTGACGTCTC TATTCTGCCC CAGCTGGAGC ATTGCAGCTC CAGGAAGATG	1901
	AATACATGGC TTGGCATTTT CTATGGTTAC AAGGGGCTGC TGCTGCTGCT GGGAATCTTC	1961
	CTTGCTTATG AGACCAAGAG TGTGTCCACT GAGAAGATCA ATGATCACCG GGCTGTGGGC	2021
30	ATGGCTATCT ACAATGTGGC AGTCCTGTGC CTCATCACTG CTCCTGTCAC CATGATTCTG	2081
	TCCAGCCAGC AGGATGCAGC CTTTGCCTTT GCCTCTTTG CCATAGTTTT CTCCTCTAT	2141
35	ATCACTCTTG TTGTGCTCTT TGTGCCCAAG ATGCGCAGGC TGATCACCCG AGGGGAATGG	2201
	CAGTCGGAGG CGCAGGACAC CATGAAGACA GGGTCATCGA CCAACAACAA CGAGGAGGAG	2261
	AAGTCCCGGC TGTTGGAGAA GGAGAACCGT GAACTGGAAA AGATCATTGC TGAGAAAGAG	2321
40	GAGCGTGTCT CTGAACTGCG CCATCAACTC CAGTCTCGGC AGCAGCTCCG CTCCCGGCGC	2381
	CACCCACCGA CACCCCCAGA ACCCTCTGGG GGCCTGCCCA GGGGACCCCC TGAGCCCCCC	2441

5	(2)		(i) 5 () (1	SEQUI A) LI B) T	ENCE ENGTI (PE:	SEQ CHAI H: 49 amir	RACTI 96 ar	ERIST	rics							
10			MOI	LECUI	LE TY	/PE: ESCR	prot	cein	SEQ :	ID NO	D: 8!	5 :				
15	Met 1	Leu	Leu	Leu	Leu 5	Leu	Leu	Ala	Pro	Leu 10	Phe	Leu	Arg	Pro	Pro 15	Gly
20	Ala	Gly	Gly	Ala 20	Gln	Thr	Pro	Asn	Ala 25	Thr	Ser	Glu	Gly	Cys 30	Gln	Ile
	Ile	His	Pro 35	Pro	Trp	Glu	Gly	Gly 40	Ile	Arg	Tyr	Arg	Gly 45	Leu	Thr	Arg
25	Asp	Gln 50	Val	Lys	Ala	Ile	Asn 55	Phe	Leu	Pro	Val	Asp 60	Tyr	Glu	Ile	Glu
	Туr 65	Val	Суѕ	Arg	Gly	Glu 70	Arg	Glu	Val	Val	Gly 75	Pro	Lys	Val	Arg	Lys 80
30	Cys	Leu	Ala	Asn	Gly 85	Ser	Trp	Thr	Asp	Met 90	Asp	Thr	Pro	Ser	Arg 95	Cys
35	Val	Arg	Ile	Cys 100	Ser	Lys	Ser	Tyr	Leu 105	Thr	Leu	Glu	Asn	Gly 110	Lys	Val
	Phe	Leu	Thr 115	Gly	Gly	Asp	Leu	Pro 120	Ala	Leu	Asp	Gly	Ala 125	Arg	Val	Asp
40	Phe	Arg 130	Cys	Asp	Pro	Asp	Phe 135	His	Leu	Cys	Asp	Pro 140	Gly	Gln	Ala	Thr

Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ilé Ile Leu

150

155

Met Pro Gly Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser Ala Thr Leu His Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly Trp Lys Lys Ile Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr Ser Thr Leu Asp Asp Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile Thr Phe Arg Gln Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Ile Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu 

Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr Asp Lys Trp Ile Val Ile Ser Arg Thr His Ser Pro Thr 

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International application No.

PCT/SE 98/01947

#### CLASSIFICATION OF SUBJECT MATTER IPC6: C07K 14/705 According to International Patent Classification (IPC) or to both national classification and IPC **B. FIELDS SEARCHED** Minimum documentation searched (classification system followed by classification symbols) IPC6: C07K Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched SE, DK, FI, NO classes as above Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Category\* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Х EMBL Databas Genbank/DDBJ, accession no. 1-6,11-13, Y11044, Grifa A. et al: "GABA (gamma-amino-butyric 16-26 acid)neurotransmission: identification and fine mapping of human GABA-B receptor gene"; Biochem. Biophys. Res. Commun. 250:240-245(1998) Nature, Volume 386, March 1997, Klemens Kaupmann et al, "Expression cloning of X 1-6,11-13, 16-26 GABAB receptors uncovers similarity to metabotropic glutamate receptors" page 239 - page 246 X WO 9746675 A1 (NOVARTIS AG), 11 December 1997 1-6,11-13, (11.12.97), page 50 - page 56, claim 4 16-26 Further documents are listed in the continuation of Box C. See patent family annex. Special categories of cited documents: later document published after the international filing date or priority date and not in conflict with the application but cited to understand "A" document defining the general state of the art which is not considered the principle or theory underlying the invention to be of particular relevance "X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive "E" erlier document but published on or after the international filing date "I." document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another ortation or other step when the document is taken alone special reason (as specified) document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination "O" document referring to an oral disclosure, use, exhibition or other being obvious to a person skilled in the art document published prior to the international filing date but later than "&" document member of the same patent family the priority date claimed Date of mailing of the international search report Date of the actual completion of the international search 2 f -03- 1999 23 March 1999 Name and mailing address of the ISA Authorized officer Swedish Patent Office Box 5055, S-102 42 STOCKHOLM Patrick Andersson Telephone No. + 46 8 782 25 00 Facsimile No. + 46 8 666 02 86

International application No.
PCT/SE 98/01947

		PC1/SE 98/0	11947
C (Continu	ation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the rele	vant passages	Relevant to claim No
Р,Х	Dialog Information Services, File 34, SciSearch, Dialog accession no. 07157158, Goei VL et al: "Human gamma-aminobutyric acid B receptor gene: Complementary DNA cloning, expression, chromosomal location and genomic organization", Biological Psychiatry, 1998, V44, N8(OCT 15), p659-6	•	1-6,11-13, 16-26
A	Brain Research Bulletin, Volume 38, No 6, 199 Robert J. Washabau et al, "GABA Receptros Dorsal Motor Nucleus of the Vagus Influen Lower Esophageal Sphincter and Gastric Fu page 587 - page 594	in the ce Feline	1-6,11-13, 16-26
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International application No.
PCT/SE 98/01947

Box I	Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)
This inter	mational search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1.	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2.	Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	rnational Searching Authority found multiple inventions in this international application, as follows:
	See next page
1.	As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
4. X	No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark	on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

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a...cmational application No.

PCT/SE 98/01947

According to PCT rule 13.2, an international application shall relate to one invention only or a group of inventions linked by one or more of the same or corresponding "special technical features", i.e. features that define a contribution which each of the inventions makes over the prior art.

The claimed invention relates to a nucleic acid sequence encoding a GABAB-receptor from either human or canine origin. A possible special technical features could have been nucleotide sequence encoding a GABAB-receptor. However a nucleotide sequence encoding a human GABAB-receptor is known from Grifa A. et. al., see the search report. Thus, the present application is considered to contain the following independent inventions: Invention 1, claims 3-4, and the parts of claims 1-2 and 16-26 relating to claims 3-4: A human

GABAB-receptor 1a, and related items,

Invention 2, claims 5-6, and the parts of claims 1-2 and 16-26 relating to claims 5-6: A human GABAB-receptor 1b, and related items.

Invention 3, claims 7-8, and the parts of claims 1-2 and 16-26 relating to claims 7-8:A human GABAB-receptor 1c, and related items,

Invention 4, claims 9-10, and the parts of claims 1-2 and 16-26 relating to claims 9-10:A human GABAB-receptor 1d, and related items,

Invention 5, claims 12-13, and the parts of claims 1, 11 and 16-26 relating to claims 12-13:A canine GABAB-receptor 1a, and related items.

Invention 6, claims 14-15, and the parts of claims 1, 11 and 16-26 relating to claims 14-15:A canine GABAB-receptor 1c, and related items.

The search has been limited to inventions 1, 2 and 5.

Information on patent family members

International application No.

02/03/99 | PCT/SE 98/01947

Patent document cited in search report Publication date Patent family member(s) Publication date

WO 9746675 A1 11/12/97 AU 2028497 A 05/01/98

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